

## SECRETED PROTEINS AND POLYNUCLEOTIDES ENCODING THEM

This application is a continuation-in-part of application Ser. No. 09/092,722, filed June 5, 1998, which claims the benefit of Ser. No. 60/086,236 (converted to a provisional application from non-provisional application Ser. No. 08/873,218), filed June 11, 1997. This application is also a continuation-in-part of application Ser. No. 09/096,287 filed June 11, 1998, which claims the benefit of Ser. No. 60/086,234 (converted to a provisional application from non-provisional application 08/873,488), filed June 12, 1997. This application is also a continuation-in-part of application Ser. No. 09/098,588 filed June 17, 1998, which claims the benefit of Ser. No. 60/XXX,XXX (converted to a provisional application from non-provisional application Ser. No. 08/878,715), filed June 19, 1997. This application is also a continuation-in-part of application Ser. No. 08/958,304 filed October 27, 1997, which claims the benefit of Ser. No. 60/092,115 (converted to a provisional application from non-provisional application Ser. No. 08/887,195, filed July 2, 1997). This application is also a continuation-in-part of application Ser. No. 09/130,189 filed August 4, 1998, which claims the benefit of Ser. No. 60/XXX,XXX (converted to a provisional application from non-provisional application Ser. No. 08/906,708), filed August 6, 1997. This application is also a continuation-in-part of application Ser. No. 09/149,633 filed September 8, 1998, which claims the benefit of Ser. No. 60/093,045 (converted to a provisional application from non-provisional application Ser. No. 08/929,007), filed September 8, 1997. This application is also a continuation-in-part of application Ser. No. 09/165,960 filed October 1, 1998, which claims the benefit of Ser. No. 60/090,100 (converted to a provisional application from non-provisional application Ser. No. 08/942,813), filed October 2, 1997. This application is also a continuation-in-part of provisional application Ser. No. 09/185,936 filed November 4, 1998, which claims the benefit of Ser. No. 60/090,111 (converted to a provisional application from non-provisional application Ser. No. 08/965,789), filed November 7, 1997. The entire content of all of the above referenced applications are incorporated by reference herein.

### FIELD OF THE INVENTION

The present invention provides novel polynucleotides and proteins encoded by such polynucleotides, along with therapeutic, diagnostic and research utilities for these polynucleotides and proteins.

### BACKGROUND OF THE INVENTION

Technology aimed at the discovery of protein factors (including e.g., cytokines, such as lymphokines, interferons, CSFs and interleukins) has matured rapidly over the

past decade. The now routine hybridization cloning and expression cloning techniques clone novel polynucleotides "directly" in the sense that they rely on information directly related to the discovered protein (i.e., partial DNA/amino acid sequence of the protein in the case of hybridization cloning; activity of the protein in the case of expression cloning). More recent "indirect" cloning techniques such as signal sequence cloning, which isolates DNA sequences based on the presence of a now well-recognized secretory leader sequence motif, as well as various PCR-based or low stringency hybridization cloning techniques, have advanced the state of the art by making available large numbers of DNA/amino acid sequences for proteins that are known to have biological activity by virtue of their secreted nature in the case of leader sequence cloning, or by virtue of the cell or tissue source in the case of PCR-based techniques. It is to these proteins and the polynucleotides encoding them that the present invention is directed.

#### SUMMARY OF THE INVENTION

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:1;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:1 from nucleotide 12 to nucleotide 800;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:1 from nucleotide 78 to nucleotide 800;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:1 from nucleotide 1 to nucleotide 547;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bh389\_11 deposited under accession number ATCC 98451;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bh389\_11 deposited under accession number ATCC 98451;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bh389\_11 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bh389\_11 deposited under accession number ATCC 98451;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:2;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:2 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:2;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

5 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:1 from nucleotide 12 to nucleotide 800; the nucleotide sequence of SEQ ID NO:1 from nucleotide 78 to nucleotide 800; the nucleotide sequence of SEQ ID NO:1 from  
10 nucleotide 1 to nucleotide 547; the nucleotide sequence of the full-length protein coding sequence of clone bh389\_11 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone bh389\_11 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert  
15 of clone bh389\_11 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:2 from amino acid 1 to amino acid 178. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of  
20 SEQ ID NO:2 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:2, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:2 having biological activity, the fragment comprising the amino acid sequence from amino acid 126 to amino acid 135 of SEQ ID NO:2.

25 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:1.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

30 (a) the amino acid sequence of SEQ ID NO:2;  
(b) the amino acid sequence of SEQ ID NO:2 from amino acid 1 to amino acid 178;

(c) fragments of the amino acid sequence of SEQ ID NO:2 comprising eight consecutive amino acids of SEQ ID NO:2; and

35 (d) the amino acid sequence encoded by the cDNA insert of clone bh389\_11 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:2 or the amino acid sequence

of SEQ ID NO:2 from amino acid 1 to amino acid 178. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:2 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:2, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:2 having biological activity, the fragment comprising the amino acid sequence from amino acid 126 to amino acid 135 of SEQ ID NO:2.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:3;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:3 from nucleotide 100 to nucleotide 882;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:3 from nucleotide 635 to nucleotide 867;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bk112\_15 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bk112\_15 deposited under accession number ATCC 98451;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bk112\_15 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bk112\_15 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:4;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:4;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:3 from nucleotide 100 to nucleotide 882; the nucleotide sequence of SEQ ID NO:3 from nucleotide 635 to nucleotide 867; the nucleotide sequence of the full-length protein



coding sequence of clone bk112\_15 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone bk112\_15 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bk112\_15 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:4 from amino acid 200 to amino acid 256. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:4, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment comprising the amino acid sequence from amino acid 125 to amino acid 134 of SEQ ID NO:4.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:3.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:4;
- (b) the amino acid sequence of SEQ ID NO:4 from amino acid 200 to amino acid 256;
- (c) fragments of the amino acid sequence of SEQ ID NO:4 comprising eight consecutive amino acids of SEQ ID NO:4; and
- (d) the amino acid sequence encoded by the cDNA insert of clone bk112\_15 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:4 or the amino acid sequence of SEQ ID NO:4 from amino acid 200 to amino acid 256. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:4, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment comprising the amino acid sequence from amino acid 125 to amino acid 134 of SEQ ID NO:4.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:5;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:5 from nucleotide 245 to nucleotide 520;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:5 from nucleotide 181 to nucleotide 527;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bk200\_13 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bk200\_13 deposited under accession number ATCC 98451;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bk200\_13 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bk200\_13 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:6;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:6;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:5 from nucleotide 245 to nucleotide 520; the nucleotide sequence of SEQ ID NO:5 from nucleotide 181 to nucleotide 527; the nucleotide sequence of the full-length protein coding sequence of clone bk200\_13 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone bk200\_13 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bk200\_13 deposited under accession number ATCC 98451. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:6, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6

having biological activity, the fragment comprising the amino acid sequence from amino acid 41 to amino acid 50 of SEQ ID NO:6.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:5.

5 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:6;
- (b) fragments of the amino acid sequence of SEQ ID NO:6 comprising eight  
10 consecutive amino acids of SEQ ID NO:6; and
- (c) the amino acid sequence encoded by the cDNA insert of clone bk200\_13 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:6. In further preferred  
15 embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:6, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6 having biological activity, the fragment  
20 comprising the amino acid sequence from amino acid 41 to amino acid 50 of SEQ ID NO:6.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:7;
- 25 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:7 from nucleotide 365 to nucleotide 784;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:7 from nucleotide 518 to nucleotide 784;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length  
30 protein coding sequence of clone di386\_3 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone di386\_3 deposited under accession number ATCC 98451;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein  
35 coding sequence of clone di386\_3 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone di386\_3 deposited under accession number ATCC 98451;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:8;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:8 having biological activity, the fragment comprising eight  
5 consecutive amino acids of SEQ ID NO:8;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

10 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:7 from nucleotide 365 to nucleotide 784; the nucleotide sequence of SEQ ID NO:7 from nucleotide 518 to nucleotide 784; the nucleotide sequence of the full-length  
15 protein coding sequence of clone di386\_3 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone di386\_3 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone di386\_3 deposited under accession number ATCC 98451. In yet other  
20 preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:8 from amino acid 1 to amino acid 140. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:8 having biological activity, the fragment preferably comprising eight  
25 (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:8, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:8 having biological activity, the fragment comprising the amino acid sequence from amino acid 65 to amino acid 74 of SEQ ID NO:8.

Other embodiments provide the gene corresponding to the cDNA sequence of  
30 SEQ ID NO:7 or SEQ ID NO:9.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:8;
- 35 (b) the amino acid sequence of SEQ ID NO:8 from amino acid 1 to amino acid 140;
- (c) fragments of the amino acid sequence of SEQ ID NO:8 comprising eight consecutive amino acids of SEQ ID NO:8; and

(d) the amino acid sequence encoded by the cDNA insert of clone di386\_3 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:8 or the amino acid sequence of SEQ ID NO:8 from amino acid 1 to amino acid 140. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:8 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:8, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:8 having biological activity, the fragment comprising the amino acid sequence from amino acid 65 to amino acid 74 of SEQ ID NO:8.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:10;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:10 from nucleotide 191 to nucleotide 781;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:10 from nucleotide 56 to nucleotide 492;
- 20 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone em397\_2 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone em397\_2 deposited under accession number ATCC 98451;
- 25 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone em397\_2 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone em397\_2 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:11;
- 30 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:11 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:11;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- 35 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:10 from nucleotide 191 to nucleotide 781; the nucleotide sequence of SEQ ID NO:10 from nucleotide 56 to nucleotide 492; the nucleotide sequence of the full-length protein coding sequence of clone em397\_2 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone em397\_2 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone em397\_2 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:11 from amino acid 1 to amino acid 101. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:11 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:11, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:11 having biological activity, the fragment comprising the amino acid sequence from amino acid 93 to amino acid 102 of SEQ ID NO:11.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:10.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:11;
- (b) the amino acid sequence of SEQ ID NO:11 from amino acid 1 to amino acid 101;
- (c) fragments of the amino acid sequence of SEQ ID NO:11 comprising eight consecutive amino acids of SEQ ID NO:11; and
- (d) the amino acid sequence encoded by the cDNA insert of clone em397\_2 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:11 or the amino acid sequence of SEQ ID NO:11 from amino acid 1 to amino acid 101. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:11 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:11, or a protein comprising a fragment of the

amino acid sequence of SEQ ID NO:11 having biological activity, the fragment comprising the amino acid sequence from amino acid 93 to amino acid 102 of SEQ ID NO:11.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:12;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:12 from nucleotide 65 to nucleotide 1636;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:12 from nucleotide 482 to nucleotide 1636;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:12 from nucleotide 487 to nucleotide 1006;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fh170\_7 deposited under accession number ATCC 98451;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fh170\_7 deposited under accession number ATCC 98451;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fh170\_7 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fh170\_7 deposited under accession number ATCC 98451;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:13;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:13 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:13;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:12 from nucleotide 65 to nucleotide 1636; the nucleotide sequence of SEQ ID NO:12 from nucleotide 482 to nucleotide 1636; the nucleotide sequence of SEQ ID NO:12 from nucleotide 487 to nucleotide 1006; the nucleotide sequence of the full-length protein coding sequence of clone fh170\_7 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone fh170\_7

deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fh170\_7 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:13 from amino acid 142 to amino acid 314. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:13 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:13, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:13 having biological activity, the fragment comprising the amino acid sequence from amino acid 257 to amino acid 266 of SEQ ID NO:13.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:12.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:13;
- (b) the amino acid sequence of SEQ ID NO:13 from amino acid 142 to amino acid 314;
- (c) fragments of the amino acid sequence of SEQ ID NO:13 comprising eight consecutive amino acids of SEQ ID NO:13; and
- (d) the amino acid sequence encoded by the cDNA insert of clone fh170\_7 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:13 or the amino acid sequence of SEQ ID NO:13 from amino acid 142 to amino acid 314. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:13 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:13, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:13 having biological activity, the fragment comprising the amino acid sequence from amino acid 257 to amino acid 266 of SEQ ID NO:13.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:15;



(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:15 from nucleotide 41 to nucleotide 550;

(c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fn53\_4 deposited under accession number ATCC 98451;

(d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fn53\_4 deposited under accession number ATCC 98451;

(e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fn53\_4 deposited under accession number ATCC 98451;

(f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fn53\_4 deposited under accession number ATCC 98451;

(g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:16;

(h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:16 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:16;

(i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;

(j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above; and

(k) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:15 from nucleotide 41 to nucleotide 550; the nucleotide sequence of the full-length protein coding sequence of clone fn53\_4 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone fn53\_4 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fn53\_4 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:16 from amino acid 40 to amino acid 170. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:16 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:16, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:16 having biological activity, the fragment comprising the amino acid sequence from amino acid 80 to amino acid 89 of SEQ ID NO:16.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:15, SEQ ID NO:14 or SEQ ID NO:17 .

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
5 from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:16;
- (b) the amino acid sequence of SEQ ID NO:16 from amino acid 40 to amino acid 170;
- (c) fragments of the amino acid sequence of SEQ ID NO:16 comprising eight  
10 consecutive amino acids of SEQ ID NO:16; and
- (d) the amino acid sequence encoded by the cDNA insert of clone fn53\_4 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:16 or the amino acid  
15 sequence of SEQ ID NO:16 from amino acid 40 to amino acid 170. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:16 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:16, or a protein comprising a fragment of the  
20 amino acid sequence of SEQ ID NO:16 having biological activity, the fragment comprising the amino acid sequence from amino acid 80 to amino acid 89 of SEQ ID NO:16.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 25 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:18;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:18 from nucleotide 84 to nucleotide 404;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:18 from nucleotide 78 to nucleotide 493;
- 30 (d) a polynucleotide comprising the nucleotide sequence of the full- length protein coding sequence of clone fq505\_4 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fq505\_4 deposited under accession number ATCC 98451;
- 35 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fq505\_4 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fq505\_4 deposited under accession number ATCC 98451;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:19;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:19 having biological activity, the fragment comprising  
5 eight consecutive amino acids of SEQ ID NO:19;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

10 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:18 from nucleotide 84 to nucleotide 404; the nucleotide sequence of SEQ ID NO:18 from nucleotide 78 to nucleotide 493; the nucleotide sequence of the full-length protein  
15 coding sequence of clone fq505\_4 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone fq505\_4 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fq505\_4 deposited under accession number ATCC 98451. In yet other  
20 preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:19 from amino acid 23 to amino acid 107. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:19 having biological activity, the fragment preferably comprising eight  
25 (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:19, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:19 having biological activity, the fragment comprising the amino acid sequence from amino acid 48 to amino acid 57 of SEQ ID NO:19.

Other embodiments provide the gene corresponding to the cDNA sequence of  
30 SEQ ID NO:18.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:19;
- 35 (b) the amino acid sequence of SEQ ID NO:19 from amino acid 23 to amino acid 107;
- (c) fragments of the amino acid sequence of SEQ ID NO:19 comprising eight consecutive amino acids of SEQ ID NO:19; and

(d) the amino acid sequence encoded by the cDNA insert of clone fq505\_4 deposited under accession number ATCC 98451; the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:19 or the amino acid sequence of SEQ ID NO:19 from amino acid 23 to amino acid 107. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:19 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:19, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:19 having biological activity, the fragment comprising the amino acid sequence from amino acid 48 to amino acid 57 of SEQ ID NO:19.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:20;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:20 from nucleotide 1439 to nucleotide 1744;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:20 from nucleotide 1241 to nucleotide 1754;
- 20 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fw13\_9 deposited under accession number ATCC 98451;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fw13\_9 deposited under accession number ATCC 98451;
- 25 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fw13\_9 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fw13\_9 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:21;
- 30 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:21 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:21;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- 35 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:20 from nucleotide 1439 to nucleotide 1744; the nucleotide sequence of SEQ ID NO:20 from nucleotide 1241 to nucleotide 1754; the nucleotide sequence of the full-length protein coding sequence of clone fw13\_9 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone fw13\_9 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fw13\_9 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:21 from amino acid 1 to amino acid 57. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:21 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:21, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:21 having biological activity, the fragment comprising the amino acid sequence from amino acid 46 to amino acid 55 of SEQ ID NO:21.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:20.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:21;
- (b) the amino acid sequence of SEQ ID NO:21 from amino acid 1 to amino acid 57;
- (c) fragments of the amino acid sequence of SEQ ID NO:21 comprising eight consecutive amino acids of SEQ ID NO:21; and
- (d) the amino acid sequence encoded by the cDNA insert of clone fw13\_9 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:21 or the amino acid sequence of SEQ ID NO:21 from amino acid 1 to amino acid 57. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:21 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty)

consecutive amino acids of SEQ ID NO:21, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:21 having biological activity, the fragment comprising the amino acid sequence from amino acid 46 to amino acid 55 of SEQ ID NO:21.

5 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:22;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:22 from nucleotide 47 to nucleotide 919;
- 10 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:22 from nucleotide 124 to nucleotide 452;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone gg619\_2 deposited under accession number ATCC 98451;
- 15 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone gg619\_2 deposited under accession number ATCC 98451;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone gg619\_2 deposited under accession number ATCC 98451;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert
- 20 of clone gg619\_2 deposited under accession number ATCC 98451;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:23;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:23 having biological activity, the fragment comprising
- 25 eight consecutive amino acids of SEQ ID NO:23;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- 30 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:22 from nucleotide 47 to nucleotide 919; the nucleotide sequence of SEQ ID NO:22 from nucleotide 124 to nucleotide 452; the nucleotide sequence of the full-length protein

35 coding sequence of clone gg619\_2 deposited under accession number ATCC 98451; or the nucleotide sequence of a mature protein coding sequence of clone gg619\_2 deposited under accession number ATCC 98451. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert

of clone gg619\_2 deposited under accession number ATCC 98451. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:23 from amino acid 27 to amino acid 135. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:23 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:23, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:23 having biological activity, the fragment comprising the amino acid sequence from amino acid 140 to amino acid 149 of SEQ ID NO:23.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:22.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:23;
- (b) the amino acid sequence of SEQ ID NO:23 from amino acid 27 to amino acid 135;
- (c) fragments of the amino acid sequence of SEQ ID NO:23 comprising eight consecutive amino acids of SEQ ID NO:23; and
- (d) the amino acid sequence encoded by the cDNA insert of clone gg619\_2 deposited under accession number ATCC 98451;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:23 or the amino acid sequence of SEQ ID NO:23 from amino acid 27 to amino acid 135. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:23 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:23, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:23 having biological activity, the fragment comprising the amino acid sequence from amino acid 140 to amino acid 149 of SEQ ID NO:23.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:35;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:35 from nucleotide 2178 to nucleotide 2513;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:35 from nucleotide 2364 to nucleotide 2513;

(d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:35 from nucleotide 1980 to nucleotide 2311;

5 (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cl181\_3 deposited under accession number ATCC 98456;

(f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cl181\_3 deposited under accession number ATCC 98456;

10 (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cl181\_3 deposited under accession number ATCC 98456;

(h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cl181\_3 deposited under accession number ATCC 98456;

15 (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:36;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:36 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:36;

20 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

25 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:35 from nucleotide 2178 to nucleotide 2513; the nucleotide sequence of SEQ ID NO:35 from nucleotide 2364 to nucleotide 2513; the nucleotide sequence of SEQ ID NO:35 from nucleotide 1980 to nucleotide 2311; the nucleotide sequence of the full-length protein coding sequence of clone cl181\_3 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone  
30 cl181\_3 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cl181\_3 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide  
35 encoding a protein comprising the amino acid sequence of SEQ ID NO:36 from amino acid 1 to amino acid 67. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:36 having biological activity, the fragment preferably



comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:36, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:36 having biological activity, the fragment comprising the amino acid sequence from amino acid 51 to amino acid 60 of SEQ ID  
 5 NO:36.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:35.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
 10 from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:36;
- (b) the amino acid sequence of SEQ ID NO:36 from amino acid 1 to amino acid 67;
- (c) fragments of the amino acid sequence of SEQ ID NO:36 comprising eight  
 15 consecutive amino acids of SEQ ID NO:36; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cl181\_3 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:36 or the amino acid  
 20 sequence of SEQ ID NO:36 from amino acid 1 to amino acid 67. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:36 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:36, or a protein comprising a fragment of the  
 25 amino acid sequence of SEQ ID NO:36 having biological activity, the fragment comprising the amino acid sequence from amino acid 51 to amino acid 60 of SEQ ID NO:36.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 30 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:37;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:37 from nucleotide 207 to nucleotide 893;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:37 from nucleotide 1 to nucleotide 527;
- 35 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cr1044\_1 deposited under accession number ATCC 98456;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cr1044\_1 deposited under accession number ATCC 98456;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cr1044\_1 deposited under accession number ATCC 98456;

5 (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cr1044\_1 deposited under accession number ATCC 98456;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:38;

10 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:38 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:38;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

15 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:37 from nucleotide 207 to nucleotide 893; the nucleotide sequence of SEQ ID NO:37 from nucleotide 1 to nucleotide 527; the nucleotide sequence of the full-length protein coding sequence of clone cr1044\_1 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone cr1044\_1 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cr1044\_1 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:38 from amino acid 1 to amino acid 107. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:38 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:38, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:38 having biological activity, the fragment comprising the amino acid sequence from amino acid 109 to amino acid 118 of SEQ ID NO:38.

35 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:37.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:38;
  - 5 (b) the amino acid sequence of SEQ ID NO:38 from amino acid 1 to amino acid 107;
  - (c) fragments of the amino acid sequence of SEQ ID NO:38 comprising eight consecutive amino acids of SEQ ID NO:38; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone cr1044\_1
  - 10 deposited under accession number ATCC 98456;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:38 or the amino acid sequence of SEQ ID NO:38 from amino acid 1 to amino acid 107. In further preferred
- 15 amino acid sequence of SEQ ID NO:38 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:38, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:38 having biological activity, the fragment comprising the amino acid sequence from amino acid 109 to amino acid 118 of SEQ ID
- 20 NO:38.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:39;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:39
- 25 from nucleotide 77 to nucleotide 400;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:39 from nucleotide 118 to nucleotide 392;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cz251\_1 deposited under accession number ATCC
- 30 98456;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cz251\_1 deposited under accession number ATCC 98456;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cz251\_1 deposited under accession number ATCC 98456;
- 35 (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cz251\_1 deposited under accession number ATCC 98456;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:40;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:40 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:40;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

10 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:39 from nucleotide 77 to nucleotide 400; the nucleotide sequence of SEQ ID NO:39 from nucleotide 118 to nucleotide 392; the nucleotide sequence of the full-length protein coding sequence of clone cz251\_1 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone cz251\_1  
15 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cz251\_1 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:40 from amino acid 15 to  
20 amino acid 105. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:40 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:40, or a polynucleotide encoding a protein comprising a fragment of the amino acid  
25 sequence of SEQ ID NO:40 having biological activity, the fragment comprising the amino acid sequence from amino acid 49 to amino acid 58 of SEQ ID NO:40.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:39.

30 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:40;

(b) the amino acid sequence of SEQ ID NO:40 from amino acid 15 to amino acid 105;

35 (c) fragments of the amino acid sequence of SEQ ID NO:40 comprising eight consecutive amino acids of SEQ ID NO:40; and

(d) the amino acid sequence encoded by the cDNA insert of clone cz251\_1 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:40 or the amino acid sequence of SEQ ID NO:40 from amino acid 15 to amino acid 105. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:40 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:40, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:40 having biological activity, the fragment comprising the amino acid sequence from amino acid 49 to amino acid 58 of SEQ ID NO:40.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:41;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:41 from nucleotide 13 to nucleotide 501;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:41 from nucleotide 1 to nucleotide 506;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone dd12\_7 deposited under accession number ATCC 98456;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone dd12\_7 deposited under accession number ATCC 98456;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone dd12\_7 deposited under accession number ATCC 98456;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone dd12\_7 deposited under accession number ATCC 98456;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:42;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:42 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:42;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:41 from nucleotide 13 to nucleotide 501; the nucleotide sequence of SEQ ID NO:41 from nucleotide 1 to nucleotide 506; the nucleotide sequence of the full-length protein coding sequence of clone dd12\_7 deposited under accession number ATCC 98456; or  
 5 the nucleotide sequence of a mature protein coding sequence of clone dd12\_7 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone dd12\_7 deposited under accession number ATCC 98456. In further preferred  
 10 embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:42 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:42, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:42 having biological activity, the fragment comprising the amino acid sequence from  
 15 amino acid 76 to amino acid 85 of SEQ ID NO:42.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:41.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
 20 from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:42;
- (b) fragments of the amino acid sequence of SEQ ID NO:42 comprising eight consecutive amino acids of SEQ ID NO:42; and
- (c) the amino acid sequence encoded by the cDNA insert of clone dd12\_7  
 25 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:42. In further preferred  
 30 embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:42 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:42, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:42 having biological activity, the fragment comprising the amino acid sequence from amino acid 76 to amino acid 85 of SEQ ID NO:42.

35 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:43;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:43 from nucleotide 778 to nucleotide 1083;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:9 from nucleotide 931 to nucleotide 1083;

5 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:43 from nucleotide 802 to nucleotide 1056;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fn191\_3 deposited under accession number ATCC 98456;

10 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fn191\_3 deposited under accession number ATCC 98456;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fn191\_3 deposited under accession number ATCC 98456;

15 (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fn191\_3 deposited under accession number ATCC 98456;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:44;

20 (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:44 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:44;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

25 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:43 from nucleotide 778 to nucleotide 1083; the nucleotide sequence of SEQ ID NO:43 from nucleotide 931 to nucleotide 1083; the nucleotide sequence of SEQ ID NO:43 from nucleotide 802 to nucleotide 1056; the nucleotide sequence of the full-length protein coding sequence of clone fn191\_3 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone fn191\_3 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fn191\_3 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:10 from amino acid 1 to amino acid 93. In further preferred embodiments, the present invention

provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:44 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:44, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:44 having biological activity, the fragment comprising the amino acid sequence from amino acid 46 to amino acid 55 of SEQ ID NO:44.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:43.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:43;
- (b) the amino acid sequence of SEQ ID NO:43 from amino acid 1 to amino acid 93;
- (c) fragments of the amino acid sequence of SEQ ID NO:43 comprising eight consecutive amino acids of SEQ ID NO:43; and
- (d) the amino acid sequence encoded by the cDNA insert of clone fn191\_3 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:44 or the amino acid sequence of SEQ ID NO:44 from amino acid 1 to amino acid 93. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:44 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:44, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:44 having biological activity, the fragment comprising the amino acid sequence from amino acid 46 to amino acid 55 of SEQ ID NO:44.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:45;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:45 from nucleotide 390 to nucleotide 1355;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:45 from nucleotide 1384 to nucleotide 1736;



(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone gm196\_4 deposited under accession number ATCC 98456;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone gm196\_4 deposited under accession number ATCC 98456;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone gm196\_4 deposited under accession number ATCC 98456;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone gm196\_4 deposited under accession number ATCC 98456;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:46;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:46 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:46;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:45 from nucleotide 390 to nucleotide 1355; the nucleotide sequence of SEQ ID NO:45 from nucleotide 1384 to nucleotide 1736; the nucleotide sequence of the full-length protein coding sequence of clone gm196\_4 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone gm196\_4 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone gm196\_4 deposited under accession number ATCC 98456. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:46 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:46, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:46 having biological activity, the fragment comprising the amino acid sequence from amino acid 156 to amino acid 165 of SEQ ID NO:46.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:45.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:46;
  - 5 (b) fragments of the amino acid sequence of SEQ ID NO:46 comprising eight consecutive amino acids of SEQ ID NO:46; and
  - (c) the amino acid sequence encoded by the cDNA insert of clone gm196\_4 deposited under accession number ATCC 98456;
- the protein being substantially free from other mammalian proteins. Preferably such
- 10 protein comprises the amino acid sequence of SEQ ID NO:46. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:46 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty)
- 15 amino acid sequence of SEQ ID NO:46 having biological activity, the fragment comprising the amino acid sequence from amino acid 156 to amino acid 165 of SEQ ID NO:46.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 20 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:47;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:47 from nucleotide 879 to nucleotide 1391;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:47 from nucleotide 519 to nucleotide 1074;
- 25 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone gn114\_1 deposited under accession number ATCC 98456;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone gn114\_1 deposited under accession number ATCC 98456;
- 30 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone gn114\_1 deposited under accession number ATCC 98456;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone gn114\_1 deposited under accession number ATCC 98456;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence
- 35 of SEQ ID NO:48;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:48 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:48;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

5 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:47 from nucleotide 879 to nucleotide 1391; the nucleotide sequence of SEQ ID NO:47 from nucleotide 519 to nucleotide 1074; the nucleotide sequence of the full-length protein coding sequence of clone gn114\_1 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone gn114\_1 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone gn114\_1 deposited under accession number ATCC 98456.

10 In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:48 from amino acid 1 to amino acid 65. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:48 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:48, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:48 having biological activity, the fragment comprising the amino acid sequence from amino acid 80 to amino acid 89 of SEQ ID NO:48.

25 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:47.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- 30 (a) the amino acid sequence of SEQ ID NO:48;
- (b) the amino acid sequence of SEQ ID NO:48 from amino acid 1 to amino acid 65;
- (c) fragments of the amino acid sequence of SEQ ID NO:48 comprising eight consecutive amino acids of SEQ ID NO:48; and
- 35 (d) the amino acid sequence encoded by the cDNA insert of clone gn114\_1 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:48 or the amino acid

sequence of SEQ ID NO:48 from amino acid 1 to amino acid 65. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:48 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:48, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:48 having biological activity, the fragment comprising the amino acid sequence from amino acid 80 to amino acid 89 of SEQ ID NO:48.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:49;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:49 from nucleotide 225 to nucleotide 1508;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:49 from nucleotide 252 to nucleotide 1508;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:49 from nucleotide 1 to nucleotide 302;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone hj968\_2 deposited under accession number ATCC 98456;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone hj968\_2 deposited under accession number ATCC 98456;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone hj968\_2 deposited under accession number ATCC 98456;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone hj968\_2 deposited under accession number ATCC 98456;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:50;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:50 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:50;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:49 from nucleotide 225 to nucleotide 1508; the nucleotide sequence of SEQ ID NO:49 from nucleotide 252 to nucleotide 1508; the nucleotide sequence of SEQ ID NO:49 from nucleotide 1 to nucleotide 302; the nucleotide sequence of the full-length protein coding sequence of clone hj968\_2 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone hj968\_2 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone hj968\_2 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:50 from amino acid 1 to amino acid 26. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:50 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:50, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:50 having biological activity, the fragment comprising the amino acid sequence from amino acid 209 to amino acid 218 of SEQ ID NO:50.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:50.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:50;
- (b) the amino acid sequence of SEQ ID NO:50 from amino acid 1 to amino acid 26;
- (c) fragments of the amino acid sequence of SEQ ID NO:50 comprising eight consecutive amino acids of SEQ ID NO:50; and
- (d) the amino acid sequence encoded by the cDNA insert of clone hj968\_2 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:50 or the amino acid sequence of SEQ ID NO:50 from amino acid 1 to amino acid 26. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:50 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:50, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:50 having biological activity, the fragment

comprising the amino acid sequence from amino acid 209 to amino acid 218 of SEQ ID NO:50.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 5 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:51;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:51 from nucleotide 1113 to nucleotide 1274;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:51 from nucleotide 1233 to nucleotide 1274;
- 10 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:51 from nucleotide 894 to nucleotide 1309;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone hk10\_3 deposited under accession number ATCC 98456;
- 15 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone hk10\_3 deposited under accession number ATCC 98456;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone hk10\_3 deposited under accession number ATCC 98456;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
20 of clone hk10\_3 deposited under accession number ATCC 98456;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:52;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:52 having biological activity, the fragment comprising  
25 eight consecutive amino acids of SEQ ID NO:52;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- 30 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:51 from nucleotide 1113 to nucleotide 1274; the nucleotide sequence of SEQ ID NO:51 from nucleotide 1233 to nucleotide 1274; the nucleotide sequence of SEQ ID  
35 NO:51 from nucleotide 894 to nucleotide 1309; the nucleotide sequence of the full-length protein coding sequence of clone hk10\_3 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone hk10\_3 deposited under accession number ATCC 98456. In other preferred

embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone hk10\_3 deposited under accession number ATCC 98456. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:52 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:52, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:52 having biological activity, the fragment comprising the amino acid sequence from amino acid 22 to amino acid 31 of SEQ ID NO:52.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:51.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:52;
- (b) fragments of the amino acid sequence of SEQ ID NO:52 comprising eight consecutive amino acids of SEQ ID NO:52; and
- (c) the amino acid sequence encoded by the cDNA insert of clone hk10\_3 deposited under accession number ATCC 98456;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:52. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:52 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:52, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:52 having biological activity, the fragment comprising the amino acid sequence from amino acid 22 to amino acid 31 of SEQ ID NO:52.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:53;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:53 from nucleotide 96 to nucleotide 1145;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:53 from nucleotide 109 to nucleotide 539;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone hm236\_1 deposited under accession number ATCC 98456;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone hm236\_1 deposited under accession number ATCC 98456;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone hm236\_1 deposited under accession number ATCC 98456;

5 (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone hm236\_1 deposited under accession number ATCC 98456;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:54;

10 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:54 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:54;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

15 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:54 from nucleotide 96 to nucleotide 1145; the nucleotide sequence of SEQ ID NO:54 from nucleotide 109 to nucleotide 539; the nucleotide sequence of the full-length protein coding sequence of clone hm236\_1 deposited under accession number ATCC 98456; or the nucleotide sequence of a mature protein coding sequence of clone hm236\_1 deposited under accession number ATCC 98456. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone hm236\_1 deposited under accession number ATCC 98456. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:54 from amino acid 6 to amino acid 148. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:54 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:54, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:54 having biological activity, the fragment comprising the amino acid sequence from amino acid 170 to amino acid 179 of SEQ ID NO:54.

35 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:53.



In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:54;
  - 5 (b) the amino acid sequence of SEQ ID NO:54 from amino acid 6 to amino acid 148;
  - (c) fragments of the amino acid sequence of SEQ ID NO:54 comprising eight consecutive amino acids of SEQ ID NO:54; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone hm236\_1
  - 10 deposited under accession number ATCC 98456;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:54 or the amino acid sequence of SEQ ID NO:43 from amino acid 6 to amino acid 148. In further preferred
- 15 amino acid sequence of SEQ ID NO:54 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:54, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:54 having biological activity, the fragment comprising the amino acid sequence from amino acid 170 to amino acid 179 of SEQ ID
- 20 NO:54.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:67;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:67
- 25 from nucleotide 185 to nucleotide 1600;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:67 from nucleotide 1403 to nucleotide 1600;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:67 from nucleotide 1 to nucleotide 850;
- 30 (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone do15\_4 deposited under accession number ATCC 98468;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone do15\_4 deposited under accession number ATCC 98468;
- 35 (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone do15\_4 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone do15\_4 deposited under accession number ATCC 98468;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:68;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:68 having biological activity, the fragment comprising  
5 eight consecutive amino acids of SEQ ID NO:68;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

10 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:67 from nucleotide 185 to nucleotide 1600; the nucleotide sequence of SEQ ID NO:67 from nucleotide 1403 to nucleotide 1600; the nucleotide sequence of SEQ ID  
15 NO:67 from nucleotide 1 to nucleotide 850; the nucleotide sequence of the full-length protein coding sequence of clone do15\_4 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone do15\_4 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert  
20 of clone do15\_4 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:2 from amino acid 1 to amino acid 222. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of  
25 SEQ ID NO:68 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:68, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:68 having biological activity, the fragment comprising the amino acid sequence from amino acid 231 to amino acid 240 of SEQ ID NO:68.

30 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:67.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

35 (a) the amino acid sequence of SEQ ID NO:68;

(b) the amino acid sequence of SEQ ID NO:68 from amino acid 1 to amino acid 222;

(c) fragments of the amino acid sequence of SEQ ID NO:68 comprising eight consecutive amino acids of SEQ ID NO:68; and

(d) the amino acid sequence encoded by the cDNA insert of clone do15\_4 deposited under accession number ATCC 98468;

5 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:68 or the amino acid sequence of SEQ ID NO:68 from amino acid 1 to amino acid 222. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:68 having biological activity, the fragment  
10 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:68, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:68 having biological activity, the fragment comprising the amino acid sequence from amino acid 231 to amino acid 240 of SEQ ID NO:68.

15 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:69;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:69 from nucleotide 47 to nucleotide 2065;

20 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:69 from nucleotide 1086 to nucleotide 1848;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone dx290\_1 deposited under accession number ATCC 98468;

25 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone dx290\_1 deposited under accession number ATCC 98468;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone dx290\_1 deposited under accession number ATCC 98468;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert  
30 of clone dx290\_1 deposited under accession number ATCC 98468;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:70;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:70 having biological activity, the fragment comprising  
35 eight consecutive amino acids of SEQ ID NO:70;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:69 from nucleotide 47 to nucleotide 2065; the nucleotide sequence of SEQ ID NO:69 from nucleotide 1086 to nucleotide 1848; the nucleotide sequence of the full-length protein coding sequence of clone dx290\_1 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone dx290\_1  
10 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone dx290\_1 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:70 from amino acid 312 to  
15 amino acid 600. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:70 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:70, or a polynucleotide encoding a protein comprising a fragment of the amino acid  
20 sequence of SEQ ID NO:70 having biological activity, the fragment comprising the amino acid sequence from amino acid 331 to amino acid 340 of SEQ ID NO:70.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:70.

In other embodiments, the present invention provides a composition  
25 comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:70;
- (b) the amino acid sequence of SEQ ID NO:70 from amino acid 312 to amino acid 600;
- 30 (c) fragments of the amino acid sequence of SEQ ID NO:70 comprising eight consecutive amino acids of SEQ ID NO:70; and

- (d) the amino acid sequence encoded by the cDNA insert of clone dx290\_1 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such  
35 protein comprises the amino acid sequence of SEQ ID NO:70 or the amino acid sequence of SEQ ID NO:4 from amino acid 312 to amino acid 600. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:70 having biological activity, the fragment

preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:70, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:70 having biological activity, the fragment comprising the amino acid sequence from amino acid 331 to amino acid 340 of SEQ ID  
 5 NO:70.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:71;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:71  
 10 from nucleotide 107 to nucleotide 724;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:71 from nucleotide 218 to nucleotide 724;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:71 from nucleotide 536 to nucleotide 866;
- 15 (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ek390\_4 deposited under accession number ATCC 98468;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ek390\_4 deposited under accession number ATCC 98468;
- 20 (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ek390\_4 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ek390\_4 deposited under accession number ATCC 98468;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence  
 25 of SEQ ID NO:72;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:72 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:72;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h)  
 30 above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

35 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:71 from nucleotide 107 to nucleotide 724; the nucleotide sequence of SEQ ID NO:71 from nucleotide 218 to nucleotide 724; the nucleotide sequence of SEQ ID NO:71 from nucleotide 536 to nucleotide 866; the nucleotide sequence of the full-length protein

coding sequence of clone ek390\_4 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone ek390\_4 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ek390\_4 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:6 from amino acid 6 to amino acid 92. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:72 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:72, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:72 having biological activity, the fragment comprising the amino acid sequence from amino acid 97 to amino acid 106 of SEQ ID NO:72.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:71.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:72;
- (b) the amino acid sequence of SEQ ID NO:6 from amino acid 6 to amino acid 92;
- (c) fragments of the amino acid sequence of SEQ ID NO:72 comprising eight consecutive amino acids of SEQ ID NO:72; and
- (d) the amino acid sequence encoded by the cDNA insert of clone ek390\_4 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:72 or the amino acid sequence of SEQ ID NO:72 from amino acid 6 to amino acid 92. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:72 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:72, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:72 having biological activity, the fragment comprising the amino acid sequence from amino acid 97 to amino acid 106 of SEQ ID NO:72.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:73;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:73 from nucleotide 31 to nucleotide 1230;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:73 from nucleotide 289 to nucleotide 1230;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:73 from nucleotide 344 to nucleotide 1119;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone er471\_7 deposited under accession number ATCC 98468;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone er471\_7 deposited under accession number ATCC 98468;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone er471\_7 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone er471\_7 deposited under accession number ATCC 98468;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:74;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:74 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:74;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:73 from nucleotide 31 to nucleotide 1230; the nucleotide sequence of SEQ ID NO:73 from nucleotide 289 to nucleotide 1230; the nucleotide sequence of SEQ ID NO:73 from nucleotide 344 to nucleotide 1119; the nucleotide sequence of the full-length protein coding sequence of clone er471\_7 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone er471\_7 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone er471\_7 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:74 from amino acid 111 to

amino acid 363. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:74 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:74, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:74 having biological activity, the fragment comprising the amino acid sequence from amino acid 195 to amino acid 204 of SEQ ID NO:74.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:73.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO51;
- (b) the amino acid sequence of SEQ ID NO:74 from amino acid 111 to amino acid 363;
- (c) fragments of the amino acid sequence of SEQ ID NO:8 comprising eight consecutive amino acids of SEQ ID NO:74; and
- (d) the amino acid sequence encoded by the cDNA insert of clone er471\_7 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:74 or the amino acid sequence of SEQ ID NO:74 from amino acid 111 to amino acid 363. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:74 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:74, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:74 having biological activity, the fragment comprising the amino acid sequence from amino acid 195 to amino acid 204 of SEQ ID NO:74.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:75;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:75 from nucleotide 62 to nucleotide 322;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:75 from nucleotide 571 to nucleotide 878;



(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fs40\_3 deposited under accession number ATCC 98468;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fs40\_3 deposited under accession number ATCC 98468;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone fs40\_3 deposited under accession number ATCC 98468;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone fs40\_3 deposited under accession number ATCC 98468;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:76;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:76 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:76;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:75 from nucleotide 62 to nucleotide 322; the nucleotide sequence of SEQ ID NO:75 from nucleotide 571 to nucleotide 878; the nucleotide sequence of the full-length protein coding sequence of clone fs40\_3 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone fs40\_3 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone fs40\_3 deposited under accession number ATCC 98468. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:76 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:76, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:76 having biological activity, the fragment comprising the amino acid sequence from amino acid 38 to amino acid 47 of SEQ ID NO:76.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:75.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:76;
  - 5 (b) fragments of the amino acid sequence of SEQ ID NO:76 comprising eight consecutive amino acids of SEQ ID NO:76; and
  - (c) the amino acid sequence encoded by the cDNA insert of clone fs40\_3 deposited under accession number ATCC 98468;
- the protein being substantially free from other mammalian proteins. Preferably such
- 10 protein comprises the amino acid sequence of SEQ ID NO:76. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:76 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:76, or a protein comprising a fragment of the
- 15 amino acid sequence of SEQ ID NO:76 having biological activity, the fragment comprising the amino acid sequence from amino acid 38 to amino acid 47 of SEQ ID NO:76.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 20 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:77;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:77 from nucleotide 43 to nucleotide 1671;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:77 from nucleotide 112 to nucleotide 1671;
- 25 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:77 from nucleotide 224 to nucleotide 679;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ga63\_6 deposited under accession number ATCC 98468;
- 30 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ga63\_6 deposited under accession number ATCC 98468;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ga63\_6 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert
- 35 of clone ga63\_6 deposited under accession number ATCC 98468;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:78;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:78 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:78;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

10 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:77 from nucleotide 43 to nucleotide 1671; the nucleotide sequence of SEQ ID NO:77 from nucleotide 112 to nucleotide 1671; the nucleotide sequence of SEQ ID NO:77 from nucleotide 224 to nucleotide 679; the nucleotide sequence of the full-length protein coding sequence of clone ga63\_6 deposited under accession number ATCC 98468; or  
15 the nucleotide sequence of a mature protein coding sequence of clone ga63\_6 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ga63\_6 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein  
20 comprising the amino acid sequence of SEQ ID NO:78 from amino acid 62 to amino acid 212. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:78 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID  
25 NO:78, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:78 having biological activity, the fragment comprising the amino acid sequence from amino acid 266 to amino acid 275 of SEQ ID NO:78.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:77.

30 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:78;
- (b) the amino acid sequence of SEQ ID NO:78 from amino acid 62 to amino  
35 acid 212;
- (c) fragments of the amino acid sequence of SEQ ID NO:78 comprising eight consecutive amino acids of SEQ ID NO:12; and

(d) the amino acid sequence encoded by the cDNA insert of clone ga63\_6 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:78 or the amino acid  
 5 sequence of SEQ ID NO:78 from amino acid 62 to amino acid 212. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:78 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:78, or a protein comprising a fragment of the  
 10 amino acid sequence of SEQ ID NO:78 having biological activity, the fragment comprising the amino acid sequence from amino acid 266 to amino acid 275 of SEQ ID NO:78.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:79;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:79 from nucleotide 17 to nucleotide 523;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:79 from nucleotide 77 to nucleotide 523;
- 20 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:79 from nucleotide 1 to nucleotide 392;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone gm335\_4 deposited under accession number ATCC 98468;
- 25 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone gm335\_4 deposited under accession number ATCC 98468;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone gm335\_4 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
 30 of clone gm335\_4 deposited under accession number ATCC 98468;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:80;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:80 having biological activity, the fragment comprising  
 35 eight consecutive amino acids of SEQ ID NO:80;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:79 from nucleotide 17 to nucleotide 523; the nucleotide sequence of SEQ ID NO:79 from nucleotide 77 to nucleotide 523; the nucleotide sequence of SEQ ID NO:79 from nucleotide 1 to nucleotide 392; the nucleotide sequence of the full-length protein coding sequence of clone gm335\_4 deposited under accession number ATCC 98468; or the  
10 nucleotide sequence of a mature protein coding sequence of clone gm335\_4 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone gm335\_4 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a  
15 protein comprising the amino acid sequence of SEQ ID NO:80 from amino acid 1 to amino acid 125. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:80 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID  
20 NO:80, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:80 having biological activity, the fragment comprising the amino acid sequence from amino acid 79 to amino acid 88 of SEQ ID NO:80.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:79.

25 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:80;
- (b) the amino acid sequence of SEQ ID NO:80 from amino acid 1 to amino  
30 acid 125;
- (c) fragments of the amino acid sequence of SEQ ID NO:80 comprising eight consecutive amino acids of SEQ ID NO:80; and
- (d) the amino acid sequence encoded by the cDNA insert of clone gm335\_4 deposited under accession number ATCC 98468;

35 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:80 or the amino acid sequence of SEQ ID NO:80 from amino acid 1 to amino acid 125. In further preferred embodiments, the present invention provides a protein comprising a fragment of the

amino acid sequence of SEQ ID NO:80 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:80, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:80 having biological activity, the fragment comprising the amino acid sequence from amino acid 79 to amino acid 88 of SEQ ID NO:80.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:81;
- 10 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:81 from nucleotide 2 to nucleotide 991;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:81 from nucleotide 62 to nucleotide 991;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:81
- 15 from nucleotide 2 to nucleotide 504;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone hy370\_9 deposited under accession number ATCC 98468;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA
- 20 insert of clone hy370\_9 deposited under accession number ATCC 98468;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone hy370\_9 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone hy370\_9 deposited under accession number ATCC 98468;
- 25 (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:82;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:82 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:82;
- 30 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one
- 35 of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:81 from nucleotide 2 to nucleotide 991; the nucleotide sequence of SEQ ID NO:81 from nucleotide 62 to nucleotide 991; the nucleotide sequence of SEQ ID NO:81 from

nucleotide 2 to nucleotide 504; the nucleotide sequence of the full-length protein coding sequence of clone hy370\_9 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone hy370\_9 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone hy370\_9 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:82 from amino acid 1 to amino acid 167. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:82 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:82, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:82 having biological activity, the fragment comprising the amino acid sequence from amino acid 160 to amino acid 169 of SEQ ID NO:82.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:81.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:82;
- (b) the amino acid sequence of SEQ ID NO:82 from amino acid 1 to amino acid 167;
- (c) fragments of the amino acid sequence of SEQ ID NO:82 comprising eight consecutive amino acids of SEQ ID NO:82; and
- (d) the amino acid sequence encoded by the cDNA insert of clone hy370\_9 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:82 or the amino acid sequence of SEQ ID NO:82 from amino acid 1 to amino acid 167. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:82 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:82, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:82 having biological activity, the fragment comprising the amino acid sequence from amino acid 160 to amino acid 169 of SEQ ID NO:82.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:83;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:83  
5 from nucleotide 77 to nucleotide 616;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:83  
from nucleotide 164 to nucleotide 616;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:83  
from nucleotide 1 to nucleotide 415;
- 10 (e) a polynucleotide comprising the nucleotide sequence of the full-length  
protein coding sequence of clone ie47\_4 deposited under accession number ATCC  
98468;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA  
insert of clone ie47\_4 deposited under accession number ATCC 98468;
- 15 (g) a polynucleotide comprising the nucleotide sequence of a mature protein  
coding sequence of clone ie47\_4 deposited under accession number ATCC 98468;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
of clone ie47\_4 deposited under accession number ATCC 98468;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence  
20 of SEQ ID NO:84;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino  
acid sequence of SEQ ID NO:84 having biological activity, the fragment comprising  
eight consecutive amino acids of SEQ ID NO:84;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h)  
25 above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i)  
or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one  
of the polynucleotides specified in (a)-(j).

30 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID  
NO:83 from nucleotide 77 to nucleotide 616; the nucleotide sequence of SEQ ID NO:83  
from nucleotide 164 to nucleotide 616; the nucleotide sequence of SEQ ID NO:83 from  
nucleotide 1 to nucleotide 415; the nucleotide sequence of the full-length protein coding  
sequence of clone ie47\_4 deposited under accession number ATCC 98468; or the  
35 nucleotide sequence of a mature protein coding sequence of clone ie47\_4 deposited  
under accession number ATCC 98468. In other preferred embodiments, the  
polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert  
of clone ie47\_4 deposited under accession number ATCC 98468. In yet other preferred



embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:84 from amino acid 1 to amino acid 113. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of  
 5 SEQ ID NO:84 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:84, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:84 having biological activity, the fragment comprising the amino acid sequence from amino acid 85 to amino acid 94 of SEQ ID NO:84.

10 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:83.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- 15 (a) the amino acid sequence of SEQ ID NO:84;
- (b) the amino acid sequence of SEQ ID NO:84 from amino acid 1 to amino acid 113;
- (c) fragments of the amino acid sequence of SEQ ID NO:84 comprising eight consecutive amino acids of SEQ ID NO:84; and
- 20 (d) the amino acid sequence encoded by the cDNA insert of clone ie47\_4 deposited under accession number ATCC 98468;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:84 or the amino acid sequence of SEQ ID NO:84 from amino acid 1 to amino acid 113. In further preferred  
 25 embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:84 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:84, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:84 having biological activity, the fragment  
 30 comprising the amino acid sequence from amino acid 85 to amino acid 94 of SEQ ID NO:84.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:85;
- 35 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:85 from nucleotide 564 to nucleotide 2813;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:85 from nucleotide 705 to nucleotide 2813;

(d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:85 from nucleotide 793 to nucleotide 1628;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone s195\_10 deposited under accession number ATCC 98468;

(f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone s195\_10 deposited under accession number ATCC 98468;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone s195\_10 deposited under accession number ATCC 98468;

(h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone s195\_10 deposited under accession number ATCC 98468;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:86;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:86 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:86;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:85 from nucleotide 564 to nucleotide 2813; the nucleotide sequence of SEQ ID NO:85 from nucleotide 705 to nucleotide 2813; the nucleotide sequence of SEQ ID NO:85 from nucleotide 793 to nucleotide 1628; the nucleotide sequence of the full-length protein coding sequence of clone s195\_10 deposited under accession number ATCC 98468; or the nucleotide sequence of a mature protein coding sequence of clone s195\_10 deposited under accession number ATCC 98468. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone s195\_10 deposited under accession number ATCC 98468. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:86 from amino acid 78 to amino acid 355. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:86 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:86, or a polynucleotide encoding a protein comprising a fragment

of the amino acid sequence of SEQ ID NO:86 having biological activity, the fragment comprising the amino acid sequence from amino acid 370 to amino acid 379 of SEQ ID NO:86.

Other embodiments provide the gene corresponding to the cDNA sequence of  
5 SEQ ID NO:85.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:86;
  - 10 (b) the amino acid sequence of SEQ ID NO:86 from amino acid 78 to amino acid 355;
  - (c) fragments of the amino acid sequence of SEQ ID NO:86 comprising eight consecutive amino acids of SEQ ID NO:86; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone s195\_10  
15 deposited under accession number ATCC 98468;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:86 or the amino acid sequence of SEQ ID NO:86 from amino acid 78 to amino acid 355. In further preferred  
20 amino acid sequence of SEQ ID NO:86 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:86, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:86 having biological activity, the fragment comprising the amino acid sequence from amino acid 370 to amino acid 379 of SEQ ID  
25 NO:86.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:97;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:97  
30 from nucleotide 516 to nucleotide 797;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:97 from nucleotide 606 to nucleotide 797;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:97 from nucleotide 1 to nucleotide 773;
- 35 (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bf228\_14 deposited under accession number ATCC 98482;

(f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bf228\_14 deposited under accession number ATCC 98482;

(g) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone bf228\_14 deposited under accession number ATCC 98482;

(h) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone bf228\_14 deposited under accession number ATCC 98482;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:98;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:98 having biological activity;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:97 from nucleotide 516 to nucleotide 797; the nucleotide sequence of SEQ ID NO:97 from nucleotide 606 to nucleotide 797; the nucleotide sequence of SEQ ID NO:97 from nucleotide 1 to nucleotide 773; the nucleotide sequence of the full-length protein coding sequence of clone bf228\_14 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone bf228\_14 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone bf228\_14 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:98 from amino acid 1 to amino acid 86.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:97.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:98;

(b) the amino acid sequence of SEQ ID NO:98 from amino acid 1 to amino acid 86;

(c) fragments of the amino acid sequence of SEQ ID NO:98; and

(d) the amino acid sequence encoded by the cDNA insert of clone bf228\_14 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:98 or the amino acid

5 sequence of SEQ ID NO:98 from amino acid 1 to amino acid 86.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:99;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:99  
10 from nucleotide 137 to nucleotide 1240;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:99 from nucleotide 1 to nucleotide 1153;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bg249\_1 deposited under accession number ATCC  
15 98482;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bg249\_1 deposited under accession number ATCC 98482;

(f) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone bg249\_1 deposited under accession number ATCC  
20 98482;

(g) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone bg249\_1 deposited under accession number ATCC 98482;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:100;

25 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:100 having biological activity;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of  
30 (h) or (i) above ; and

(l) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:99 from nucleotide 137 to nucleotide 1240; the nucleotide sequence of SEQ ID  
35 NO:99 from nucleotide 1 to nucleotide 1153; the nucleotide sequence of the full-length protein coding sequence of clone bg249\_1 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone bg249\_1 deposited under accession number ATCC 98482. In other preferred

embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone bg249\_1 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:100 from amino acid 1 to amino acid 339.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:99.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:100;
- (b) the amino acid sequence of SEQ ID NO:100 from amino acid 1 to amino acid 339;
- (c) fragments of the amino acid sequence of SEQ ID NO:100; and
- (d) the amino acid sequence encoded by the cDNA insert of clone bg249\_1 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:100 or the amino acid sequence of SEQ ID NO:100 from amino acid 1 to amino acid 339.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:101;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:101 from nucleotide 26 to nucleotide 301;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:101 from nucleotide 104 to nucleotide 301;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:5 from nucleotide 1 to nucleotide 119;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bv286\_1 deposited under accession number ATCC 98482;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bv286\_1 deposited under accession number ATCC 98482;
- (g) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone bv286\_1 deposited under accession number ATCC 98482;
- (h) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone bv286\_1 deposited under accession number ATCC 98482;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:102;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:102 having biological activity;

5 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

10 (m) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:101 from nucleotide 26 to nucleotide 301; the nucleotide sequence of SEQ ID NO:101 from nucleotide 104 to nucleotide 301; the nucleotide sequence of SEQ ID NO:101 from nucleotide 1 to nucleotide 119; the nucleotide sequence of the full-length  
15 protein coding sequence of clone bv286\_1 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone bv286\_1 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone bv286\_1 deposited under accession number ATCC 98482. In  
20 yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:102 from amino acid 1 to amino acid 31.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:101.

25 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:102;
- (b) the amino acid sequence of SEQ ID NO:102 from amino acid 1 to amino  
30 acid 31;
- (c) fragments of the amino acid sequence of SEQ ID NO:102; and
- (d) the amino acid sequence encoded by the cDNA insert of clone bv286\_1 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such  
35 protein comprises the amino acid sequence of SEQ ID NO:102 or the amino acid sequence of SEQ ID NO:102 from amino acid 1 to amino acid 31.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:103;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:103 from nucleotide 663 to nucleotide 755;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:103 from nucleotide 1 to nucleotide 850;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone co36\_1 deposited under accession number ATCC 98482;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone co36\_1 deposited under accession number ATCC 98482;
- (f) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone co36\_1 deposited under accession number ATCC 98482;
- (g) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone co36\_1 deposited under accession number ATCC 98482;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:104;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:104 having biological activity;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:103 from nucleotide 663 to nucleotide 755; the nucleotide sequence of SEQ ID NO:103 from nucleotide 1 to nucleotide 850; the nucleotide sequence of the full-length protein coding sequence of clone co36\_1 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone co36\_1 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone co36\_1 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:104 from amino acid 1 to amino acid 22.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:103.



In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:104;
- 5 (b) the amino acid sequence of SEQ ID NO:8 from amino acid 1 to amino acid 22;
- (c) fragments of the amino acid sequence of SEQ ID NO:104; and
- (d) the amino acid sequence encoded by the cDNA insert of clone co36\_1 deposited under accession number ATCC 98482;
- 10 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:104 or the amino acid sequence of SEQ ID NO:104 from amino acid 1 to amino acid 22.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:105;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:105 from nucleotide 127 to nucleotide 783;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:105 from nucleotide 172 to nucleotide 783;
- 20 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:105 from nucleotide 7 to nucleotide 462;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cp116\_1 deposited under accession number ATCC 98482;
- 25 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cp116\_1 deposited under accession number ATCC 98482;
- (g) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone cp116\_1 deposited under accession number ATCC 98482;
- 30 (h) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone cp116\_1 deposited under accession number ATCC 98482;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:106;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:106 having biological activity;
- 35 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(j).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:105 from nucleotide 127 to nucleotide 783; the nucleotide sequence of SEQ ID NO:105 from nucleotide 172 to nucleotide 783; the nucleotide sequence of SEQ ID NO:105 from nucleotide 7 to nucleotide 462; the nucleotide sequence of the full-length protein coding sequence of clone cp116\_1 deposited under accession number ATCC  
10 98482; or the nucleotide sequence of the mature protein coding sequence of clone cp116\_1 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone cp116\_1 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide  
15 encoding a protein comprising the amino acid sequence of SEQ ID NO:106 from amino acid 1 to amino acid 112.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:105.

20 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:106;
- (b) the amino acid sequence of SEQ ID NO:106 from amino acid 1 to amino acid 112;
- 25 (c) fragments of the amino acid sequence of SEQ ID NO:106; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cp116\_1 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:106 or the amino acid  
30 sequence of SEQ ID NO:106 from amino acid 1 to amino acid 112.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:108;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:108  
35 from nucleotide 231 to nucleotide 533;
- (c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cw1195\_2 deposited under accession number ATCC 98482;

(d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cw1195\_2 deposited under accession number ATCC 98482;

(e) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone cw1195\_2 deposited under accession number ATCC 98482;

(f) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone cw1195\_2 deposited under accession number ATCC 98482;

(g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:109;

(h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:109 having biological activity;

(i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;

(j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above ; and

(k) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO75 from nucleotide 231 to nucleotide 533; the nucleotide sequence of the full-length protein coding sequence of clone cw1195\_2 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone cw1195\_2 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone cw1195\_2 deposited under accession number ATCC 98482.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:108, SEQ ID NO74 or SEQ ID NO:110 .

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:109;

(b) fragments of the amino acid sequence of SEQ ID NO:109; and

(c) the amino acid sequence encoded by the cDNA insert of clone cw1195\_2 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:109.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:111;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:111 from nucleotide 645 to nucleotide 782;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:111 from nucleotide 10 to nucleotide 773;

5 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone fh13\_10 deposited under accession number ATCC 98482;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone fh13\_10 deposited under accession number ATCC 98482;

10 (f) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone fh13\_10 deposited under accession number ATCC 98482;

(g) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone fh13\_10 deposited under accession number ATCC 98482;

15 (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:112;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:112 having biological activity;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

20 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(i).

25 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:111 from nucleotide 645 to nucleotide 782; the nucleotide sequence of SEQ ID NO:111 from nucleotide 10 to nucleotide 773; the nucleotide sequence of the full-length protein coding sequence of clone fh13\_10 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone  
30 fh13\_10 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone fh13\_10 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:112 from amino  
35 acid 1 to amino acid 43.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:111.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:112;
- 5 (b) the amino acid sequence of SEQ ID NO:16 from amino acid 1 to amino acid 43;
- (c) fragments of the amino acid sequence of SEQ ID NO:112; and
- (d) the amino acid sequence encoded by the cDNA insert of clone fh13\_10 deposited under accession number ATCC 98482;
- 10 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:112 or the amino acid sequence of SEQ ID NO:112 from amino acid 1 to amino acid 43.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:113;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:113 from nucleotide 94 to nucleotide 216;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:17 from nucleotide 160 to nucleotide 216;
- 20 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:113 from nucleotide 20 to nucleotide 193;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone gc57\_4 deposited under accession number ATCC 98482;
- 25 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone gc57\_4 deposited under accession number ATCC 98482;
- (g) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone gc57\_4 deposited under accession number ATCC 98482;
- 30 (h) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone gc57\_4 deposited under accession number ATCC 98482;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:114;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:114 having biological activity;
- 35 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(j).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:113 from nucleotide 94 to nucleotide 216; the nucleotide sequence of SEQ ID NO:113 from nucleotide 160 to nucleotide 216; the nucleotide sequence of SEQ ID NO:113 from nucleotide 20 to nucleotide 193; the nucleotide sequence of the full-length protein coding sequence of clone gc57\_4 deposited under accession number ATCC  
10 98482; or the nucleotide sequence of the mature protein coding sequence of clone gc57\_4 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone gc57\_4 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide  
15 encoding a protein comprising the amino acid sequence of SEQ ID NO:18 from amino acid 1 to amino acid 33.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:113.

20 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:114;
- (b) the amino acid sequence of SEQ ID NO:114 from amino acid 1 to amino acid 33;
- 25 (c) fragments of the amino acid sequence of SEQ ID NO:114; and
- (d) the amino acid sequence encoded by the cDNA insert of clone gc57\_4 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:114 or the amino acid  
30 sequence of SEQ ID NO:114 from amino acid 1 to amino acid 33.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:115;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:115  
35 from nucleotide 2 to nucleotide 943;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:115 from nucleotide 2 to nucleotide 670;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone h1165\_3 deposited under accession number ATCC 98482;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone h1165\_3 deposited under accession number ATCC 98482;

(f) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone h1165\_3 deposited under accession number ATCC 98482;

(g) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone h1165\_3 deposited under accession number ATCC 98482;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:116;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:116 having biological activity;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:115 from nucleotide 2 to nucleotide 943; the nucleotide sequence of SEQ ID NO:115 from nucleotide 2 to nucleotide 670; the nucleotide sequence of the full-length protein coding sequence of clone h1165\_3 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone h1165\_3 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone h1165\_3 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:116 from amino acid 1 to amino acid 223.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:115.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:116;

(b) the amino acid sequence of SEQ ID NO:116 from amino acid 1 to amino acid 223;

(c) fragments of the amino acid sequence of SEQ ID NO:116; and

(d) the amino acid sequence encoded by the cDNA insert of clone h1165\_3 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:116 or the amino acid sequence of SEQ ID NO:116 from amino acid 1 to amino acid 223.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:117;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:117 from nucleotide 1242 to nucleotide 1457;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:21 from nucleotide 1326 to nucleotide 1457;

(d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:117 from nucleotide 869 to nucleotide 1544;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone hb752\_1 deposited under accession number ATCC 98482;

(f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone hb752\_1 deposited under accession number ATCC 98482;

(g) a polynucleotide comprising the nucleotide sequence of the mature protein coding sequence of clone hb752\_1 deposited under accession number ATCC 98482;

(h) a polynucleotide encoding the mature protein encoded by the cDNA insert of clone hb752\_1 deposited under accession number ATCC 98482;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:118;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:118 having biological activity;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(j).



Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:117 from nucleotide 1242 to nucleotide 1457; the nucleotide sequence of SEQ ID NO:117 from nucleotide 1326 to nucleotide 1457; the nucleotide sequence of SEQ ID NO:117 from nucleotide 869 to nucleotide 1544; the nucleotide sequence of the full-length protein coding sequence of clone hb752\_1 deposited under accession number ATCC 98482; or the nucleotide sequence of the mature protein coding sequence of clone hb752\_1 deposited under accession number ATCC 98482. In other preferred embodiments, the polynucleotide encodes the full-length or mature protein encoded by the cDNA insert of clone hb752\_1 deposited under accession number ATCC 98482. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:118 from amino acid 1 to amino acid 69.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:117.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:118;
- (b) the amino acid sequence of SEQ ID NO:118 from amino acid 1 to amino acid 69;
- (c) fragments of the amino acid sequence of SEQ ID NO:118; and
- (d) the amino acid sequence encoded by the cDNA insert of clone hb752\_1 deposited under accession number ATCC 98482;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:118 or the amino acid sequence of SEQ ID NO:118 from amino acid 1 to amino acid 69.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:129;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:129 from nucleotide 864 to nucleotide 1340;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:129 from nucleotide 1 to nucleotide 1175;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bi127\_5 deposited under accession number ATCC 98501;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bi127\_5 deposited under accession number ATCC 98501;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bi127\_5 deposited under accession number ATCC 98501;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bi127\_5 deposited under accession number ATCC 98501;

5 (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:130;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:130 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:130;

10 (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one  
15 of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:129 from nucleotide 864 to nucleotide 1340; the nucleotide sequence of SEQ ID NO:129 from nucleotide 1 to nucleotide 1175; the nucleotide sequence of the full-length protein coding sequence of clone bi127\_5 deposited under accession number ATCC  
20 98501; or the nucleotide sequence of a mature protein coding sequence of clone bi127\_5 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bi127\_5 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a  
25 protein comprising the amino acid sequence of SEQ ID NO:130 from amino acid 1 to amino acid 104. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:130 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID  
30 NO:130, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:130 having biological activity, the fragment comprising the amino acid sequence from amino acid 74 to amino acid 83 of SEQ ID NO:130.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:129.

35 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:130;

(b) the amino acid sequence of SEQ ID NO:130 from amino acid 1 to amino acid 104;

(c) fragments of the amino acid sequence of SEQ ID NO:130 comprising eight consecutive amino acids of SEQ ID NO:130; and

5 (d) the amino acid sequence encoded by the cDNA insert of clone bi127\_5 deposited under accession number ATCC 98501;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:130 or the amino acid sequence of SEQ ID NO:130 from amino acid 1 to amino acid 104. In further preferred  
10 embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:130 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:130, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:130 having biological activity, the fragment  
15 comprising the amino acid sequence from amino acid 74 to amino acid 83 of SEQ ID NO:130.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:131;

20 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:131 from nucleotide 46 to nucleotide 738;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:131 from nucleotide 346 to nucleotide 738;

(d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:3  
25 from nucleotide 688 to nucleotide 1425;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bl194\_2 deposited under accession number ATCC 98501;

(f) a polynucleotide encoding the full-length protein encoded by the cDNA  
30 insert of clone bl194\_2 deposited under accession number ATCC 98501;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bl194\_2 deposited under accession number ATCC 98501;

(h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bl194\_2 deposited under accession number ATCC 98501;

35 (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:132;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:132 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:132;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

10 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:131 from nucleotide 46 to nucleotide 738; the nucleotide sequence of SEQ ID NO:131 from nucleotide 346 to nucleotide 738; the nucleotide sequence of SEQ ID NO:131 from nucleotide 688 to nucleotide 1425; the nucleotide sequence of the full-length protein coding sequence of clone bl194\_2 deposited under accession number  
15 ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone bl194\_2 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bl194\_2 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide  
20 encoding a protein comprising the amino acid sequence of SEQ ID NO:4 from amino acid 1 to amino acid 171. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:132 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino  
25 acids of SEQ ID NO:132, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:132 having biological activity, the fragment comprising the amino acid sequence from amino acid 110 to amino acid 119 of SEQ ID NO:132.

Other embodiments provide the gene corresponding to the cDNA sequence of  
30 SEQ ID NO:131.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:132;
- 35 (b) the amino acid sequence of SEQ ID NO:132 from amino acid 1 to amino acid 171;
- (c) fragments of the amino acid sequence of SEQ ID NO:132 comprising eight consecutive amino acids of SEQ ID NO:132; and

(d) the amino acid sequence encoded by the cDNA insert of clone bl194\_2 deposited under accession number ATCC 98501;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:132 or the amino acid  
 5 sequence of SEQ ID NO:132 from amino acid 1 to amino acid 171. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:132 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:132, or a protein comprising a fragment of the  
 10 amino acid sequence of SEQ ID NO:132 having biological activity, the fragment comprising the amino acid sequence from amino acid 110 to amino acid 119 of SEQ ID NO:132.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:133;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:133 from nucleotide 234 to nucleotide 1235;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:133 from nucleotide 291 to nucleotide 1235;
- 20 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:133 from nucleotide 209 to nucleotide 1050;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cc130\_1 deposited under accession number ATCC 98501;
- 25 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cc130\_1 deposited under accession number ATCC 98501;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cc130\_1 deposited under accession number ATCC 98501;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
 30 of clone cc130\_1 deposited under accession number ATCC 98501;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:134;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:134 having biological activity, the fragment comprising  
 35 eight consecutive amino acids of SEQ ID NO:134;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

(m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:133 from nucleotide 234 to nucleotide 1235; the nucleotide sequence of SEQ ID NO:133 from nucleotide 291 to nucleotide 1235; the nucleotide sequence of SEQ ID NO:133 from nucleotide 209 to nucleotide 1050; the nucleotide sequence of the full-length protein coding sequence of clone cc130\_1 deposited under accession number  
 10 ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone cc130\_1 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cc130\_1 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide  
 15 encoding a protein comprising the amino acid sequence of SEQ ID NO:6 from amino acid 1 to amino acid 272. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:134 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino  
 20 acids of SEQ ID NO:134, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:134 having biological activity, the fragment comprising the amino acid sequence from amino acid 162 to amino acid 171 of SEQ ID NO:134.

Other embodiments provide the gene corresponding to the cDNA sequence of  
 25 SEQ ID NO:133.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:134;
  - 30 (b) the amino acid sequence of SEQ ID NO:134 from amino acid 1 to amino acid 272;
  - (c) fragments of the amino acid sequence of SEQ ID NO:134 comprising eight consecutive amino acids of SEQ ID NO:134; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone cc130\_1  
 35 deposited under accession number ATCC 98501;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:134 or the amino acid sequence of SEQ ID NO:134 from amino acid 1 to amino acid 272. In further preferred

embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:134 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:134, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:134 having biological activity, the fragment comprising the amino acid sequence from amino acid 162 to amino acid 171 of SEQ ID NO:134.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 10 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:135;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:135 from nucleotide 1554 to nucleotide 1784;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:135 from nucleotide 1659 to nucleotide 1784;
- 15 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:135 from nucleotide 1508 to nucleotide 1865;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ch582\_1 deposited under accession number ATCC 98501;
- 20 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ch582\_1 deposited under accession number ATCC 98501;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ch582\_1 deposited under accession number ATCC 98501;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ch582\_1 deposited under accession number ATCC 98501;
- 25 (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:136;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:136 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:136;
- 30 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- 35 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:135 from nucleotide 1554 to nucleotide 1784; the nucleotide sequence of SEQ ID

NO:135 from nucleotide 1659 to nucleotide 1784; the nucleotide sequence of SEQ ID NO:135 from nucleotide 1508 to nucleotide 1865; the nucleotide sequence of the full-length protein coding sequence of clone ch582\_1 deposited under accession number ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone ch582\_1 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ch582\_1 deposited under accession number ATCC 98501. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:136 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:136, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:136 having biological activity, the fragment comprising the amino acid sequence from amino acid 33 to amino acid 42 of SEQ ID NO:136.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:135.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:136;
- (b) fragments of the amino acid sequence of SEQ ID NO:136 comprising eight consecutive amino acids of SEQ ID NO:136; and
- (c) the amino acid sequence encoded by the cDNA insert of clone ch582\_1 deposited under accession number ATCC 98501;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:136. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:136 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:136, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:136 having biological activity, the fragment comprising the amino acid sequence from amino acid 33 to amino acid 42 of SEQ ID NO:136.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:137;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:137 from nucleotide 1375 to nucleotide 1605;



(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:137 from nucleotide 1107 to nucleotide 1539;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cq294\_14 deposited under accession number ATCC 98501;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cq294\_14 deposited under accession number ATCC 98501;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cq294\_14 deposited under accession number ATCC 98501;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cq294\_14 deposited under accession number ATCC 98501;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:138;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:138 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:138;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:137 from nucleotide 1375 to nucleotide 1605; the nucleotide sequence of SEQ ID NO:137 from nucleotide 1107 to nucleotide 1539; the nucleotide sequence of the full-length protein coding sequence of clone cq294\_14 deposited under accession number ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone cq294\_14 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cq294\_14 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:138 from amino acid 1 to amino acid 55. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:138 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:138, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:138 having biological activity, the fragment

comprising the amino acid sequence from amino acid 33 to amino acid 42 of SEQ ID NO:138.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:137.

5 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:138;
- (b) the amino acid sequence of SEQ ID NO:138 from amino acid 1 to amino acid 55;
- (c) fragments of the amino acid sequence of SEQ ID NO:138 comprising eight consecutive amino acids of SEQ ID NO:138; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cq294\_14 deposited under accession number ATCC 98501;

15 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:138 or the amino acid sequence of SEQ ID NO:138 from amino acid 1 to amino acid 55. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:138 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:138, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:138 having biological activity, the fragment comprising the amino acid sequence from amino acid 33 to amino acid 42 of SEQ ID NO:138.

25 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:139;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:139 from nucleotide 66 to nucleotide 1880;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:139 from nucleotide 1 to nucleotide 581;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone dd454\_1 deposited under accession number ATCC 98501;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone dd454\_1 deposited under accession number ATCC 98501;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone dd454\_1 deposited under accession number ATCC 98501;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone dd454\_1 deposited under accession number ATCC 98501;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:140;

5 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:140 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:140;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

10 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID  
15 NO:139 from nucleotide 66 to nucleotide 1880; the nucleotide sequence of SEQ ID NO:139 from nucleotide 1 to nucleotide 581; the nucleotide sequence of the full-length protein coding sequence of clone dd454\_1 deposited under accession number ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone dd454\_1 deposited under accession number ATCC 98501. In other preferred embodiments, the  
20 polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone dd454\_1 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:140 from amino acid 1 to amino acid 172. In further preferred embodiments, the present invention provides a  
25 polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:140 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:140, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:140 having biological activity, the fragment comprising  
30 the amino acid sequence from amino acid 297 to amino acid 306 of SEQ ID NO:140.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:139.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
35 from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:140;

(b) the amino acid sequence of SEQ ID NO:140 from amino acid 1 to amino acid 172;

(c) fragments of the amino acid sequence of SEQ ID NO:140 comprising eight consecutive amino acids of SEQ ID NO:140; and

(d) the amino acid sequence encoded by the cDNA insert of clone dd454\_1 deposited under accession number ATCC 98501;

5 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:140 or the amino acid sequence of SEQ ID NO:140 from amino acid 1 to amino acid 172. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:140 having biological activity, the fragment  
10 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:140, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:140 having biological activity, the fragment comprising the amino acid sequence from amino acid 297 to amino acid 306 of SEQ ID NO:140.

15 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:141;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:141 from nucleotide 462 to nucleotide 3170;

20 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:141 from nucleotide 1188 to nucleotide 1517;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone du157\_12 deposited under accession number ATCC 98724;

25 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone du157\_12 deposited under accession number ATCC 98724;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone du157\_12 deposited under accession number ATCC 98724;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert  
30 of clone du157\_12 deposited under accession number ATCC 98724;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:142;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:142 having biological activity, the fragment comprising  
35 eight consecutive amino acids of SEQ ID NO:142;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:141 from nucleotide 462 to nucleotide 3170; the nucleotide sequence of SEQ ID NO:141 from nucleotide 1188 to nucleotide 1517; the nucleotide sequence of the full-length protein coding sequence of clone du157\_12 deposited under accession number ATCC 98724; or the nucleotide sequence of a mature protein coding sequence of clone  
10 du157\_12 deposited under accession number ATCC 98724. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone du157\_12 deposited under accession number ATCC 98724. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:142 from amino  
15 acid 251 to amino acid 352. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:142 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:142, or a polynucleotide encoding a protein comprising a fragment  
20 of the amino acid sequence of SEQ ID NO:142 having biological activity, the fragment comprising the amino acid sequence from amino acid 446 to amino acid 455 of SEQ ID NO:142.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:141.

25 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:142;
  - (b) the amino acid sequence of SEQ ID NO:142 from amino acid 251 to  
30 amino acid 352;
  - (c) fragments of the amino acid sequence of SEQ ID NO:142 comprising eight consecutive amino acids of SEQ ID NO:142; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone du157\_12 deposited under accession number ATCC 98724;
- 35 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:142 or the amino acid sequence of SEQ ID NO:142 from amino acid 251 to amino acid 352. In further preferred embodiments, the present invention provides a protein comprising a

fragment of the amino acid sequence of SEQ ID NO:142 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:142, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:142 having biological activity, the fragment  
 5 comprising the amino acid sequence from amino acid 446 to amino acid 455 of SEQ ID NO:142.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:143;
- 10 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:143 from nucleotide 865 to nucleotide 1158;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:143 from nucleotide 1108 to nucleotide 1158;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:143  
 15 from nucleotide 1 to nucleotide 764;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone du372\_1 deposited under accession number ATCC 98501;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA  
 20 insert of clone du372\_1 deposited under accession number ATCC 98501;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone du372\_1 deposited under accession number ATCC 98501;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone du372\_1 deposited under accession number ATCC 98501;
- 25 (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:144;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:144 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:144;
- 30 (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one  
 35 of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:143 from nucleotide 865 to nucleotide 1158; the nucleotide sequence of SEQ ID NO:143 from nucleotide 1108 to nucleotide 1158; the nucleotide sequence of SEQ ID

NO:143 from nucleotide 1 to nucleotide 764; the nucleotide sequence of the full-length protein coding sequence of clone du372\_1 deposited under accession number ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone du372\_1 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone du372\_1 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:144 from amino acid 69 to amino acid 98. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:144 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:144, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:144 having biological activity, the fragment comprising the amino acid sequence from amino acid 44 to amino acid 53 of SEQ ID NO:144.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:143.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:144;
- (b) the amino acid sequence of SEQ ID NO:144 from amino acid 69 to amino acid 98;
- (c) fragments of the amino acid sequence of SEQ ID NO:144 comprising eight consecutive amino acids of SEQ ID NO:144; and
- (d) the amino acid sequence encoded by the cDNA insert of clone du372\_1 deposited under accession number ATCC 98501;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:144 or the amino acid sequence of SEQ ID NO:144 from amino acid 69 to amino acid 98. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:144 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:144, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:144 having biological activity, the fragment comprising the amino acid sequence from amino acid 44 to amino acid 53 of SEQ ID NO:144.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:145;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:145  
5 from nucleotide 32 to nucleotide 586;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:145 from nucleotide 92 to nucleotide 586;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:145 from nucleotide 1 to nucleotide 481;
- 10 (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ej90\_5 deposited under accession number ATCC 98501;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ej90\_5 deposited under accession number ATCC 98501;
- 15 (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ej90\_5 deposited under accession number ATCC 98501;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ej90\_5 deposited under accession number ATCC 98501;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence  
20 of SEQ ID NO:146;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:146 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:146;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h)  
25 above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

30 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:145 from nucleotide 32 to nucleotide 586; the nucleotide sequence of SEQ ID NO:145 from nucleotide 92 to nucleotide 586; the nucleotide sequence of SEQ ID NO:145 from nucleotide 1 to nucleotide 481; the nucleotide sequence of the full-length protein coding sequence of clone ej90\_5 deposited under accession number ATCC  
35 98501; or the nucleotide sequence of a mature protein coding sequence of clone ej90\_5 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ej90\_5 deposited under accession number ATCC 98501. In yet other preferred



embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:146 from amino acid 1 to amino acid 150. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:146 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:146, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:146 having biological activity, the fragment comprising the amino acid sequence from amino acid 87 to amino acid 96 of SEQ ID NO:146.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:145.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:146;
- (b) the amino acid sequence of SEQ ID NO:146 from amino acid 1 to amino acid 150;
- (c) fragments of the amino acid sequence of SEQ ID NO:146 comprising eight consecutive amino acids of SEQ ID NO:146; and
- (d) the amino acid sequence encoded by the cDNA insert of clone ej90\_5 deposited under accession number ATCC 98501;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:146 or the amino acid sequence of SEQ ID NO:146 from amino acid 1 to amino acid 150. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:146 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:146, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:146 having biological activity, the fragment comprising the amino acid sequence from amino acid 87 to amino acid 96 of SEQ ID NO:146.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:147;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:147 from nucleotide 281 to nucleotide 1786;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:147 from nucleotide 332 to nucleotide 1786;

(d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:147 from nucleotide 1 to nucleotide 574;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ic2\_6 deposited under accession number ATCC 98501;

5 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ic2\_6 deposited under accession number ATCC 98501;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ic2\_6 deposited under accession number ATCC 98501;

(h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
10 of clone ic2\_6 deposited under accession number ATCC 98501;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:148;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:148 having biological activity, the fragment comprising  
15 eight consecutive amino acids of SEQ ID NO:148;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

20 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:147 from nucleotide 281 to nucleotide 1786; the nucleotide sequence of SEQ ID NO:147 from nucleotide 332 to nucleotide 1786; the nucleotide sequence of SEQ ID  
25 NO:147 from nucleotide 1 to nucleotide 574; the nucleotide sequence of the full-length protein coding sequence of clone ic2\_6 deposited under accession number ATCC 98501; or the nucleotide sequence of a mature protein coding sequence of clone ic2\_6 deposited under accession number ATCC 98501. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert  
30 of clone ic2\_6 deposited under accession number ATCC 98501. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:148 from amino acid 1 to amino acid 98. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of  
35 SEQ ID NO:148 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:148, or a polynucleotide encoding a protein comprising a fragment of the amino

acid sequence of SEQ ID NO:148 having biological activity, the fragment comprising the amino acid sequence from amino acid 246 to amino acid 255 of SEQ ID NO:148.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:147.

5 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:148;
- (b) the amino acid sequence of SEQ ID NO:148 from amino acid 1 to amino  
10 acid 98;
- (c) fragments of the amino acid sequence of SEQ ID NO:148 comprising eight consecutive amino acids of SEQ ID NO:148; and

- (d) the amino acid sequence encoded by the cDNA insert of clone ic2\_6 deposited under accession number ATCC 98501;
- 15 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:148 or the amino acid sequence of SEQ ID NO:148 from amino acid 1 to amino acid 98. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:148 having biological activity, the fragment  
20 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:148, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:148 having biological activity, the fragment comprising the amino acid sequence from amino acid 246 to amino acid 255 of SEQ ID NO:148.

25 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:159;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:159 from nucleotide 69 to nucleotide 908;
- 30 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:159 from nucleotide 270 to nucleotide 908;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bn97\_1 deposited under accession number ATCC 98535;
- 35 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bn97\_1 deposited under accession number ATCC 98535;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bn97\_1 deposited under accession number ATCC 98535;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bn97\_1 deposited under accession number ATCC 98535;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:160;

5 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:160 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:160;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

10 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID  
15 NO:159 from nucleotide 69 to nucleotide 908; the nucleotide sequence of SEQ ID NO:159 from nucleotide 270 to nucleotide 908; the nucleotide sequence of the full-length protein coding sequence of clone bn97\_1 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone bn97\_1 deposited under accession number ATCC 98535. In other preferred  
20 embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bn97\_1 deposited under accession number ATCC 98535. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:160 from amino acid 1 to amino acid 83. In further preferred embodiments, the present invention  
25 provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:160 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:160, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:160 having biological activity, the fragment  
30 comprising the amino acid sequence from amino acid 135 to amino acid 144 of SEQ ID NO:160.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:159.

In other embodiments, the present invention provides a composition  
35 comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:160;

(b) the amino acid sequence of SEQ ID NO:160 from amino acid 1 to amino acid 83;

(c) fragments of the amino acid sequence of SEQ ID NO:160 comprising eight consecutive amino acids of SEQ ID NO:160; and

5 (d) the amino acid sequence encoded by the cDNA insert of clone bn97\_1 deposited under accession number ATCC 98535;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:160 or the amino acid sequence of SEQ ID NO:160 from amino acid 1 to amino acid 83. In further preferred  
10 embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:160 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:160, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:160 having biological activity, the fragment  
15 comprising the amino acid sequence from amino acid 135 to amino acid 144 of SEQ ID NO:160.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:161;  
20 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:161 from nucleotide 562 to nucleotide 777;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:161 from nucleotide 236 to nucleotide 673;

(d) a polynucleotide comprising the nucleotide sequence of the full-length  
25 protein coding sequence of clone bn268\_11 deposited under accession number ATCC 98535;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bn268\_11 deposited under accession number ATCC 98535;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein  
30 coding sequence of clone bn268\_11 deposited under accession number ATCC 98535;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bn268\_11 deposited under accession number ATCC 98535;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:162;

35 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:162 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:162;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

5 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:161 from nucleotide 562 to nucleotide 777; the nucleotide sequence of SEQ ID NO:161 from nucleotide 236 to nucleotide 673; the nucleotide sequence of the full-length protein coding sequence of clone bn268\_11 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone bn268\_11 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bn268\_11 deposited under accession number ATCC 98535.

15 In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:162 from amino acid 1 to amino acid 37. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:162 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:4, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:4 having biological activity, the fragment comprising the amino acid sequence from amino acid 31 to amino acid 40 of SEQ ID NO:162.

25 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:161.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- 30 (a) the amino acid sequence of SEQ ID NO:162;
- (b) the amino acid sequence of SEQ ID NO:162 from amino acid 1 to amino acid 37;
- (c) fragments of the amino acid sequence of SEQ ID NO:162 comprising eight consecutive amino acids of SEQ ID NO:162; and
- 35 (d) the amino acid sequence encoded by the cDNA insert of clone bn268\_11 deposited under accession number ATCC 98535;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:162 or the amino acid

sequence of SEQ ID NO:162 from amino acid 1 to amino acid 37. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:162 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:162, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:162 having biological activity, the fragment comprising the amino acid sequence from amino acid 31 to amino acid 40 of SEQ ID NO:162.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:163;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:163 from nucleotide 286 to nucleotide 1686;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:163 from nucleotide 544 to nucleotide 1686;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:163 from nucleotide 365 to nucleotide 1160;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cb96\_10 deposited under accession number ATCC 98535;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cb96\_10 deposited under accession number ATCC 98535;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cb96\_10 deposited under accession number ATCC 98535;
- (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cb96\_10 deposited under accession number ATCC 98535;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:164;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:164 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:164;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:163 from nucleotide 286 to nucleotide 1686; the nucleotide sequence of SEQ ID NO:163 from nucleotide 544 to nucleotide 1686; the nucleotide sequence of SEQ ID NO:163 from nucleotide 365 to nucleotide 1160; the nucleotide sequence of the full-length protein coding sequence of clone cb96\_10 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone cb96\_10 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cb96\_10 deposited under accession number ATCC 98535.

In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:164 from amino acid 28 to amino acid 395. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:164 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:164, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:164 having biological activity, the fragment comprising the amino acid sequence from amino acid 228 to amino acid 237 of SEQ ID NO:164.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:163.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:164;
- (b) the amino acid sequence of SEQ ID NO:164 from amino acid 28 to amino acid 395;
- (c) fragments of the amino acid sequence of SEQ ID NO:164 comprising eight consecutive amino acids of SEQ ID NO:164; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cb96\_10 deposited under accession number ATCC 98535;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:164 or the amino acid sequence of SEQ ID NO:164 from amino acid 28 to amino acid 395. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:164 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:164, or a protein comprising a fragment of the



amino acid sequence of SEQ ID NO:164 having biological activity, the fragment comprising the amino acid sequence from amino acid 228 to amino acid 237 of SEQ ID NO:164.

In one embodiment, the present invention provides a composition comprising  
5 an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:165;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:165 from nucleotide 99 to nucleotide 1049;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:165  
10 from nucleotide 222 to nucleotide 1049;
- (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:165 from nucleotide 632 to nucleotide 998;
- (e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cb213\_11 deposited under accession number ATCC  
15 98535;
- (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cb213\_11 deposited under accession number ATCC 98535;
- (g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cb213\_11 deposited under accession number ATCC 98535;
- 20 (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cb213\_11 deposited under accession number ATCC 98535;
- (i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:166;
- (j) a polynucleotide encoding a protein comprising a fragment of the amino  
25 acid sequence of SEQ ID NO:166 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:166;
- (k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;
- (l) a polynucleotide which encodes a species homologue of the protein of (i)  
30 or (j) above ; and
- (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:166 from nucleotide 99 to nucleotide 1049; the nucleotide sequence of SEQ ID  
35 NO:166 from nucleotide 222 to nucleotide 1049; the nucleotide sequence of SEQ ID NO:166 from nucleotide 632 to nucleotide 998; the nucleotide sequence of the full-length protein coding sequence of clone cb213\_11 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone

cb213\_11 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cb213\_11 deposited under accession number ATCC 98535. In yet other preferred embodiments, the present invention provides a polynucleotide  
 5 encoding a protein comprising the amino acid sequence of SEQ ID NO:166 from amino acid 187 to amino acid 300. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:166 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino  
 10 acids of SEQ ID NO:166, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:166 having biological activity, the fragment comprising the amino acid sequence from amino acid 153 to amino acid 162 of SEQ ID NO:166.

Other embodiments provide the gene corresponding to the cDNA sequence of  
 15 SEQ ID NO:165.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:166;
  - 20 (b) the amino acid sequence of SEQ ID NO:166 from amino acid 187 to amino acid 300;
  - (c) fragments of the amino acid sequence of SEQ ID NO:166 comprising eight consecutive amino acids of SEQ ID NO:166; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone cb213\_11  
 25 deposited under accession number ATCC 98535;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:166 or the amino acid sequence of SEQ ID NO:166 from amino acid 187 to amino acid 300. In further preferred embodiments, the present invention provides a protein comprising a  
 30 fragment of the amino acid sequence of SEQ ID NO:166 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:166, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:166 having biological activity, the fragment comprising the amino acid sequence from amino acid 153 to amino acid 162 of SEQ ID  
 35 NO:166.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:167;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:167 from nucleotide 3003 to nucleotide 3137;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:167 from nucleotide 3072 to nucleotide 3137;

5 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:167 from nucleotide 2713 to nucleotide 3114;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cj457\_4 deposited under accession number ATCC 98535;

10 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cj457\_4 deposited under accession number ATCC 98535;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cj457\_4 deposited under accession number ATCC 98535;

15 (h) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cj457\_4 deposited under accession number ATCC 98535;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:168;

20 (j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:168 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:168;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

25 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:167 from nucleotide 3003 to nucleotide 3137; the nucleotide sequence of SEQ ID NO:167 from nucleotide 3072 to nucleotide 3137; the nucleotide sequence of SEQ ID NO:167 from nucleotide 2713 to nucleotide 3114; the nucleotide sequence of the full-length protein coding sequence of clone cj457\_4 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone cj457\_4 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cj457\_4 deposited under accession number ATCC 98535. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:168 from amino acid 1 to amino acid 37. In further preferred embodiments, the present invention

provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:168 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:168, or a polynucleotide encoding a protein comprising a fragment  
 5 of the amino acid sequence of SEQ ID NO:168 having biological activity, the fragment comprising the amino acid sequence from amino acid 17 to amino acid 26 of SEQ ID NO:168.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:167.

10 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:168;
- (b) the amino acid sequence of SEQ ID NO:168 from amino acid 1 to amino  
 15 acid 37;
- (c) fragments of the amino acid sequence of SEQ ID NO:168 comprising eight consecutive amino acids of SEQ ID NO:168; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cj457\_4 deposited under accession number ATCC 98535;

20 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:168 or the amino acid sequence of SEQ ID NO:168 from amino acid 1 to amino acid 37. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:168 having biological activity, the fragment  
 25 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:168, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:168 having biological activity, the fragment comprising the amino acid sequence from amino acid 17 to amino acid 26 of SEQ ID NO:168.

30 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:169;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:169 from nucleotide 284 to nucleotide 1357;
- 35 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:169 from nucleotide 603 to nucleotide 1233;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cz653\_11 deposited under accession number ATCC 98535;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cz653\_11 deposited under accession number ATCC 98535;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cz653\_11 deposited under accession number ATCC 98535;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cz653\_11 deposited under accession number ATCC 98535;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:170;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:12 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:170;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:169 from nucleotide 284 to nucleotide 1357; the nucleotide sequence of SEQ ID NO:169 from nucleotide 603 to nucleotide 1233; the nucleotide sequence of the full-length protein coding sequence of clone cz653\_11 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone cz653\_11 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cz653\_11 deposited under accession number ATCC 98535. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:170 from amino acid 147 to amino acid 358. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:170 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:170, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:170 having biological activity, the fragment comprising the amino acid sequence from amino acid 174 to amino acid 183 of SEQ ID NO:170.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:169.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
5 from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:170;
- (b) the amino acid sequence of SEQ ID NO:170 from amino acid 147 to amino acid 358;
- (c) fragments of the amino acid sequence of SEQ ID NO:170 comprising  
10 eight consecutive amino acids of SEQ ID NO:170; and
- (d) the amino acid sequence encoded by the cDNA insert of clone cz653\_11 deposited under accession number ATCC 98535;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:170 or the amino acid  
15 sequence of SEQ ID NO:170 from amino acid 147 to amino acid 358. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:170 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:170, or a protein comprising a fragment of the  
20 amino acid sequence of SEQ ID NO:170 having biological activity, the fragment comprising the amino acid sequence from amino acid 174 to amino acid 183 of SEQ ID NO:170.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 25 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:171;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:171 from nucleotide 621 to nucleotide 1763;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:171 from nucleotide 1461 to nucleotide 1763;
- 30 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone dx138\_4 deposited under accession number ATCC 98535;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone dx138\_4 deposited under accession number ATCC 98535;
- 35 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone dx138\_4 deposited under accession number ATCC 98535;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone dx138\_4 deposited under accession number ATCC 98535;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:172;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:172 having biological activity, the fragment comprising  
5 eight consecutive amino acids of SEQ ID NO:172;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

10 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:171 from nucleotide 621 to nucleotide 1763; the nucleotide sequence of SEQ ID NO:171 from nucleotide 1461 to nucleotide 1763; the nucleotide sequence of the full-  
15 length protein coding sequence of clone dx138\_4 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone dx138\_4 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone dx138\_4 deposited under accession number ATCC 98535.  
20 In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:172 from amino acid 83 to amino acid 229. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:172 having biological activity, the fragment preferably  
25 comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:172, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:172 having biological activity, the fragment comprising the amino acid sequence from amino acid 185 to amino acid 194 of SEQ ID NO:172.

30 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:171.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

35 (a) the amino acid sequence of SEQ ID NO:172;

(b) the amino acid sequence of SEQ ID NO:172 from amino acid 83 to amino acid 229;

(c) fragments of the amino acid sequence of SEQ ID NO:172 comprising eight consecutive amino acids of SEQ ID NO:172; and

(d) the amino acid sequence encoded by the cDNA insert of clone dx138\_4 deposited under accession number ATCC 98535;

5 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:172 or the amino acid sequence of SEQ ID NO:172 from amino acid 83 to amino acid 229. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:172 having biological activity, the fragment  
10 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:172, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:172 having biological activity, the fragment comprising the amino acid sequence from amino acid 185 to amino acid 194 of SEQ ID NO:172.

15 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:173;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:173 from nucleotide 119 to nucleotide 295;

20 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:173 from nucleotide 191 to nucleotide 295;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ij167\_5 deposited under accession number ATCC 98535;

25 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ij167\_5 deposited under accession number ATCC 98535;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ij167\_5 deposited under accession number ATCC 98535;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert  
30 of clone ij167\_5 deposited under accession number ATCC 98535;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:174;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:174 having biological activity, the fragment comprising  
35 eight consecutive amino acids of SEQ ID NO:174;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;



(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:174 from nucleotide 119 to nucleotide 295; the nucleotide sequence of SEQ ID NO:174 from nucleotide 191 to nucleotide 295; the nucleotide sequence of the full-length protein coding sequence of clone ij167\_5 deposited under accession number ATCC 98535; or the nucleotide sequence of a mature protein coding sequence of clone  
10 ij167\_5 deposited under accession number ATCC 98535. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ij167\_5 deposited under accession number ATCC 98535. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:16 from amino  
15 acid 1 to amino acid 26. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:174 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:174, or a polynucleotide encoding a protein comprising a fragment  
20 of the amino acid sequence of SEQ ID NO:174 having biological activity, the fragment comprising the amino acid sequence from amino acid 24 to amino acid 33 of SEQ ID NO:174.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:174.

25 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:174;
- (b) the amino acid sequence of SEQ ID NO:174 from amino acid 1 to amino  
30 acid 26;
- (c) fragments of the amino acid sequence of SEQ ID NO:174 comprising eight consecutive amino acids of SEQ ID NO:174; and
- (d) the amino acid sequence encoded by the cDNA insert of clone ij167\_5 deposited under accession number ATCC 98535;

35 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:174 or the amino acid sequence of SEQ ID NO:174 from amino acid 1 to amino acid 26. In further preferred embodiments, the present invention provides a protein comprising a fragment of the

amino acid sequence of SEQ ID NO:174 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:174, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:174 having biological activity, the fragment comprising the amino acid sequence from amino acid 24 to amino acid 33 of SEQ ID NO:174.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:183;
- 10 (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:183 from nucleotide 25 to nucleotide 1458;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:183 from nucleotide 21 to nucleotide 730;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bd107\_16 deposited under accession number ATCC 98898;
- 15 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bd107\_16 deposited under accession number ATCC 98898;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bd107\_16 deposited under accession number ATCC 98898;
- 20 (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bd107\_16 deposited under accession number ATCC 98898;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:184;
- 25 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:184 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:184;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- 30 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:183 from nucleotide 25 to nucleotide 1458; the nucleotide sequence of SEQ ID NO:183 from nucleotide 21 to nucleotide 730; the nucleotide sequence of the full-length protein coding sequence of clone bd107\_16 deposited under accession number ATCC 98898; or the nucleotide sequence of a mature protein coding sequence of clone

bd107\_16 deposited under accession number ATCC 98898. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bd107\_16 deposited under accession number ATCC 98898. In yet other preferred embodiments, the present invention provides a polynucleotide  
5 encoding a protein comprising the amino acid sequence of SEQ ID NO:184 from amino acid 2 to amino acid 118. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:184 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino  
10 acids of SEQ ID NO:184, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:184 having biological activity, the fragment comprising the amino acid sequence from amino acid 234 to amino acid 243 of SEQ ID NO:184.

Other embodiments provide the gene corresponding to the cDNA sequence of  
15 SEQ ID NO:183.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:184;
  - 20 (b) the amino acid sequence of SEQ ID NO:184 from amino acid 2 to amino acid 118;
  - (c) fragments of the amino acid sequence of SEQ ID NO:184 comprising eight consecutive amino acids of SEQ ID NO:184; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone bd107\_16  
25 deposited under accession number ATCC 98898;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:184 or the amino acid sequence of SEQ ID NO:184 from amino acid 2 to amino acid 118. In further preferred embodiments, the present invention provides a protein comprising a fragment of the  
30 amino acid sequence of SEQ ID NO:184 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:184, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:184 having biological activity, the fragment comprising the amino acid sequence from amino acid 234 to amino acid 243 of SEQ ID  
35 NO:184.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:185;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:185 from nucleotide 6 to nucleotide 977;

(c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:185 from nucleotide 87 to nucleotide 977;

5 (d) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:185 from nucleotide 8 to nucleotide 630;

(e) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bm41\_7 deposited under accession number ATCC 98898;

10 (f) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bm41\_7 deposited under accession number ATCC 98898;

(g) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bm41\_7 deposited under accession number ATCC 98898;

(h) a polynucleotide encoding a mature protein encoded by the cDNA insert  
15 of clone bm41\_7 deposited under accession number ATCC 98898;

(i) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:186;

(j) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:186 having biological activity, the fragment comprising  
20 eight consecutive amino acids of SEQ ID NO:4;

(k) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(h) above;

(l) a polynucleotide which encodes a species homologue of the protein of (i) or (j) above ; and

25 (m) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(j).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:184 from nucleotide 6 to nucleotide 977; the nucleotide sequence of SEQ ID NO:184 from nucleotide 87 to nucleotide 977; the nucleotide sequence of SEQ ID NO:184 from  
30 nucleotide 8 to nucleotide 630; the nucleotide sequence of the full-length protein coding sequence of clone bm41\_7 deposited under accession number ATCC 98898; or the nucleotide sequence of a mature protein coding sequence of clone bm41\_7 deposited under accession number ATCC 98898. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert  
35 of clone bm41\_7 deposited under accession number ATCC 98898. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:4 from amino acid 211 to amino acid 315. In further preferred embodiments, the present invention provides a

polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:186 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:186, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:186 having biological activity, the fragment comprising the amino acid sequence from amino acid 157 to amino acid 166 of SEQ ID NO:186.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:185.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:186;
- (b) the amino acid sequence of SEQ ID NO:186 from amino acid 211 to amino acid 315;
- (c) fragments of the amino acid sequence of SEQ ID NO:186 comprising eight consecutive amino acids of SEQ ID NO:186; and
- (d) the amino acid sequence encoded by the cDNA insert of clone bm41\_7 deposited under accession number ATCC 98898;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:186 or the amino acid sequence of SEQ ID NO:4 from amino acid 211 to amino acid 315. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:186 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:186, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:186 having biological activity, the fragment comprising the amino acid sequence from amino acid 157 to amino acid 166 of SEQ ID NO:186.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:187;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:187 from nucleotide 168 to nucleotide 962;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:187 from nucleotide 351 to nucleotide 962;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone br342\_11 deposited under accession number ATCC 98551;

(e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone br342\_11 deposited under accession number ATCC 98551;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone br342\_11 deposited under accession number ATCC 98551;

5 (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone br342\_11 deposited under accession number ATCC 98551;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:188;

10 (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:188 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:188;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

15 (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:187 from nucleotide 168 to nucleotide 962; the nucleotide sequence of SEQ ID NO:187 from nucleotide 351 to nucleotide 962; the nucleotide sequence of the full-length protein coding sequence of clone br342\_11 deposited under accession number ATCC 98551; or the nucleotide sequence of a mature protein coding sequence of clone br342\_11 deposited under accession number ATCC 98551. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone br342\_11 deposited under accession number ATCC 98551. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:188 from amino acid 1 to amino acid 78. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:188 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:188, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:188 having biological activity, the fragment comprising the amino acid sequence from amino acid 127 to amino acid 136 of SEQ ID NO:188.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:187.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:188;
  - 5 (b) the amino acid sequence of SEQ ID NO:188 from amino acid 1 to amino acid 78;
  - (c) fragments of the amino acid sequence of SEQ ID NO:188 comprising eight consecutive amino acids of SEQ ID NO:188; and
  - (d) the amino acid sequence encoded by the cDNA insert of clone br342\_11
  - 10 deposited under accession number ATCC 98551;
- the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:188 or the amino acid sequence of SEQ ID NO:188 from amino acid 1 to amino acid 78. In further preferred
- 15 amino acid sequence of SEQ ID NO:188 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:188, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:188 having biological activity, the fragment comprising the amino acid sequence from amino acid 127 to amino acid 136 of SEQ ID
- 20 NO:188.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:189;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:189
- 25 from nucleotide 134 to nucleotide 493;
- (c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ej258\_11 deposited under accession number ATCC 98551;
- (d) a polynucleotide encoding the full-length protein encoded by the cDNA
- 30 insert of clone ej258\_11 deposited under accession number ATCC 98551;
- (e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ej258\_11 deposited under accession number ATCC 98551;
- (f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ej258\_11 deposited under accession number ATCC 98551;
- 35 (g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:190;

(h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:190 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:190;

(i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;

(j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above ; and

(k) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(h).

10 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:189 from nucleotide 134 to nucleotide 493; the nucleotide sequence of the full-length protein coding sequence of clone ej258\_11 deposited under accession number ATCC 98551; or the nucleotide sequence of a mature protein coding sequence of clone ej258\_11 deposited under accession number ATCC 98551. In other preferred  
15 embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ej258\_11 deposited under accession number ATCC 98551. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:190 from amino acid 1 to amino acid 64. In further preferred embodiments, the present invention  
20 provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:190 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:190, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:190 having biological activity, the fragment  
25 comprising the amino acid sequence from amino acid 55 to amino acid 64 of SEQ ID NO:190.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:189.

30 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:190;

(b) the amino acid sequence of SEQ ID NO:8 from amino acid 1 to amino acid 64;

35 (c) fragments of the amino acid sequence of SEQ ID NO:190 comprising eight consecutive amino acids of SEQ ID NO:190; and

(d) the amino acid sequence encoded by the cDNA insert of clone ej258\_11 deposited under accession number ATCC 98551;



the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:190 or the amino acid sequence of SEQ ID NO:190 from amino acid 1 to amino acid 64. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:190 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:190, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:190 having biological activity, the fragment comprising the amino acid sequence from amino acid 55 to amino acid 64 of SEQ ID NO:190.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:191;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:191 from nucleotide 14 to nucleotide 406;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:191 from nucleotide 62 to nucleotide 406;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone k232\_2x deposited under accession number ATCC 98551;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone k232\_2x deposited under accession number ATCC 98551;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone k232\_2x deposited under accession number ATCC 98551;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone k232\_2x deposited under accession number ATCC 98551;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:192;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:192 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:192;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:191 from nucleotide 14 to nucleotide 406; the nucleotide sequence of SEQ ID NO:191 from nucleotide 62 to nucleotide 406; the nucleotide sequence of the full-length protein coding sequence of clone k232\_2x deposited under accession number ATCC 98551; or the nucleotide sequence of a mature protein coding sequence of clone k232\_2x deposited under accession number ATCC 98551. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone k232\_2x deposited under accession number ATCC 98551. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:192 from amino acid 1 to amino acid 81. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:192 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:192, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:192 having biological activity, the fragment comprising the amino acid sequence from amino acid 60 to amino acid 69 of SEQ ID NO:192.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:191.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:192;
- (b) the amino acid sequence of SEQ ID NO:192 from amino acid 1 to amino acid 81;
- (c) fragments of the amino acid sequence of SEQ ID NO:192 comprising eight consecutive amino acids of SEQ ID NO:192; and
- (d) the amino acid sequence encoded by the cDNA insert of clone k232\_2x deposited under accession number ATCC 98551;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:192 or the amino acid sequence of SEQ ID NO:192 from amino acid 1 to amino acid 81. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:192 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:192, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:192 having biological activity, the fragment

comprising the amino acid sequence from amino acid 60 to amino acid 69 of SEQ ID NO:192.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 5 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:193;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:193 from nucleotide 580 to nucleotide 816;
- (c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone lf307\_5 deposited under accession number ATCC 98551;
- 10 (d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone lf307\_5 deposited under accession number ATCC 98551;
- (e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone lf307\_5 deposited under accession number ATCC 98551;
- 15 (f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone lf307\_5 deposited under accession number ATCC 98551;
- (g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:194;
- (h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:194 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:194;
- 20 (i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;
- (j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above ; and
- 25 (k) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:193 from nucleotide 580 to nucleotide 816; the nucleotide sequence of the full-length protein coding sequence of clone lf307\_5 deposited under accession number ATCC 98551; or the nucleotide sequence of a mature protein coding sequence of clone lf307\_5 deposited under accession number ATCC 98551. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone lf307\_5 deposited under accession number ATCC 98551.

35 In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:194 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:194,

or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:194 having biological activity, the fragment comprising the amino acid sequence from amino acid 34 to amino acid 43 of SEQ ID NO:194.

Other embodiments provide the gene corresponding to the cDNA sequence of  
5 SEQ ID NO:193 or SEQ ID NO:195.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:194;
- 10 (b) fragments of the amino acid sequence of SEQ ID NO:194 comprising eight consecutive amino acids of SEQ ID NO:194; and
- (c) the amino acid sequence encoded by the cDNA insert of clone lf307\_5 deposited under accession number ATCC 98551;

the protein being substantially free from other mammalian proteins. Preferably such  
15 protein comprises the amino acid sequence of SEQ ID NO:194. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:194 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:194, or a protein comprising a fragment of the  
20 amino acid sequence of SEQ ID NO:194 having biological activity, the fragment comprising the amino acid sequence from amino acid 34 to amino acid 43 of SEQ ID NO:194.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 25 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:196;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:196 from nucleotide 127 to nucleotide 627;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:196 from nucleotide 250 to nucleotide 627;
- 30 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone lr204\_1 deposited under accession number ATCC 98551;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone lr204\_1 deposited under accession number ATCC 98551;
- 35 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone lr204\_1 deposited under accession number ATCC 98551;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone lr204\_1 deposited under accession number ATCC 98551;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:197;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:197 having biological activity, the fragment comprising  
5 eight consecutive amino acids of SEQ ID NO:197;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

10 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:196 from nucleotide 127 to nucleotide 627; the nucleotide sequence of SEQ ID NO:196 from nucleotide 250 to nucleotide 627; the nucleotide sequence of the full-  
15 length protein coding sequence of clone lr204\_1 deposited under accession number ATCC 98551; or the nucleotide sequence of a mature protein coding sequence of clone lr204\_1 deposited under accession number ATCC 98551. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone lr204\_1 deposited under accession number ATCC 98551.  
20 In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:197 from amino acid 23 to amino acid 106. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:197 having biological activity, the fragment preferably  
25 comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:197, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:197 having biological activity, the fragment comprising the amino acid sequence from amino acid 78 to amino acid 87 of SEQ ID NO:197.

30 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:196.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

35 (a) the amino acid sequence of SEQ ID NO:197;

(b) the amino acid sequence of SEQ ID NO:197 from amino acid 23 to amino acid 106;

(c) fragments of the amino acid sequence of SEQ ID NO:197 comprising eight consecutive amino acids of SEQ ID NO:197; and

(d) the amino acid sequence encoded by the cDNA insert of clone Ir204\_1 deposited under accession number ATCC 98551;

5 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:197 or the amino acid sequence of SEQ ID NO:197 from amino acid 23 to amino acid 106. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:197 having biological activity, the fragment  
10 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:197, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:197 having biological activity, the fragment comprising the amino acid sequence from amino acid 78 to amino acid 87 of SEQ ID NO:197.

15 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

(a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:205;

(b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:205 from nucleotide 876 to nucleotide 1190;

20 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:205 from nucleotide 963 to nucleotide 1190;

(d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone as20\_2 deposited under accession number ATCC 98580;

25 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone as20\_2 deposited under accession number ATCC 98580;

(f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone as20\_2 deposited under accession number ATCC 98580;

(g) a polynucleotide encoding a mature protein encoded by the cDNA insert  
30 of clone as20\_2 deposited under accession number ATCC 98580;

(h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:206;

(i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:206 having biological activity, the fragment comprising  
35 eight consecutive amino acids of SEQ ID NO:206;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:37 from nucleotide 876 to nucleotide 1190; the nucleotide sequence of SEQ ID NO:205 from nucleotide 963 to nucleotide 1190; the nucleotide sequence of the full-length protein coding sequence of clone as20\_2 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone  
10 as20\_2 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone as20\_2 deposited under accession number ATCC 98580. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:2 from amino  
15 acid 1 to amino acid 60. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:206 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:206, or a polynucleotide encoding a protein comprising a fragment  
20 of the amino acid sequence of SEQ ID NO:206 having biological activity, the fragment comprising the amino acid sequence from amino acid 47 to amino acid 56 of SEQ ID NO:206.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:205.

25 A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:

(i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

30 (aa) SEQ ID NO:205, but excluding the poly(A) tail at the 3' end of SEQ ID NO:205; and

(ab) the nucleotide sequence of the cDNA insert of clone as20\_2 deposited under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

35 (iii) isolating the DNA polynucleotide detected with the probe(s);  
and

(b) a process comprising the steps of:

(i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:205, but excluding the poly(A) tail at the 3' end of SEQ ID NO:205; and

5 (bb) the nucleotide sequence of the cDNA insert of clone as20\_2 deposited under ATCC 98580;

(ii) hybridizing said primer(s) to human DNA;

(iii) amplifying human DNA sequences; and

(iv) isolating the polynucleotide product of step (b)(iii).

10 Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:205, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:205 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:1 but excluding the poly(A) tail at the 3' end of SEQ ID NO:205. In another preferred embodiment, the

15 nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:205 from nucleotide 876 to nucleotide 1190, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:205 from nucleotide 876 to nucleotide 1190, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:205 from nucleotide 876 to nucleotide 1190.

20 In a further preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:205 from nucleotide 963 to nucleotide 1190, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:205 from nucleotide 963 to nucleotide 1190, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:1 from nucleotide 963 to nucleotide 1190.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:206;

30 (b) the amino acid sequence of SEQ ID NO:206 from amino acid 1 to amino acid 60;

(c) fragments of the amino acid sequence of SEQ ID NO:206 comprising eight consecutive amino acids of SEQ ID NO:206; and

(d) the amino acid sequence encoded by the cDNA insert of clone as20\_2 deposited under accession number ATCC 98580;

35 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:206 or the amino acid sequence of SEQ ID NO:206 from amino acid 1 to amino acid 60. In further preferred



embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:206 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:206, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:206 having biological activity, the fragment comprising the amino acid sequence from amino acid 47 to amino acid 56 of SEQ ID NO:206.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 10 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:207;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:207 from nucleotide 946 to nucleotide 1095;
- (c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bf227\_8 deposited under accession number ATCC 15 98580;
- (d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bf227\_8 deposited under accession number ATCC 98580;
- (e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bf227\_8 deposited under accession number ATCC 98580;
- 20 (f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bf227\_8 deposited under accession number ATCC 98580;
- (g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:208;
- (h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:208 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:208;
- 25 (i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;
- (j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above ; and
- 30 (k) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:207 from nucleotide 946 to nucleotide 1095; the nucleotide sequence of the full-length protein coding sequence of clone bf227\_8 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone bf227\_8 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded

by the cDNA insert of clone bf227\_8 deposited under accession number ATCC 98580. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:208 from amino acid 1 to amino acid 34. In further preferred embodiments, the present invention

5 provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:208 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:208, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:208 having biological activity, the fragment

10 comprising the amino acid sequence from amino acid 19 to amino acid 28 of SEQ ID NO:208.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:207.

A further embodiment of the invention provides a process for producing an

15 isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:

(i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(aa) SEQ ID NO:207, but excluding the poly(A) tail at the 3' end of SEQ ID

20 NO:207; and

(ab) the nucleotide sequence of the cDNA insert of clone bf227\_8 deposited under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

(iii) isolating the DNA polynucleotide detected with the probe(s);

25 and

(b) a process comprising the steps of:

(i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:207, but excluding the poly(A) tail at the 3' end of SEQ ID

30 NO:207; and

(bb) the nucleotide sequence of the cDNA insert of clone bf227\_8 deposited under ATCC 98580;

(ii) hybridizing said primer(s) to human DNA;

(iii) amplifying human DNA sequences; and

35 (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:207, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:207

to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:3 but excluding the poly(A) tail at the 3' end of SEQ ID NO:207. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:207 from nucleotide 946 to nucleotide 1095, and extends contiguously  
 5 from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:207 from nucleotide 946 to nucleotide 1095, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:207 from nucleotide 946 to nucleotide 1095.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected  
 10 from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:208;
- (b) the amino acid sequence of SEQ ID NO:208 from amino acid 1 to amino acid 34;
- (c) fragments of the amino acid sequence of SEQ ID NO:208 comprising  
 15 eight consecutive amino acids of SEQ ID NO:208; and
- (d) the amino acid sequence encoded by the cDNA insert of clone bf227\_8 deposited under accession number ATCC 98580;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:208 or the amino acid  
 20 sequence of SEQ ID NO:208 from amino acid 1 to amino acid 34. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:208 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:208, or a protein comprising a fragment of the  
 25 amino acid sequence of SEQ ID NO:208 having biological activity, the fragment comprising the amino acid sequence from amino acid 19 to amino acid 28 of SEQ ID NO:208.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 30 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:209;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:209 from nucleotide 183 to nucleotide 911;
- (c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone bh157\_7 deposited under accession number ATCC  
 35 98580;
- (d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone bh157\_7 deposited under accession number ATCC 98580;

(e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone bh157\_7 deposited under accession number ATCC 98580;

(f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone bh157\_7 deposited under accession number ATCC 98580;

5 (g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:6;

(h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:6 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:6;

10 (i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;

(j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above ; and

(k) a polynucleotide that hybridizes under stringent conditions to any one  
15 of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:209 from nucleotide 183 to nucleotide 911; the nucleotide sequence of the full-length protein coding sequence of clone bh157\_7 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone  
20 bh157\_7 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone bh157\_7 deposited under accession number ATCC 98580. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:210 from amino  
25 acid 1 to amino acid 76. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:210 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:210, or a polynucleotide encoding a protein comprising a fragment  
30 of the amino acid sequence of SEQ ID NO:210 having biological activity, the fragment comprising the amino acid sequence from amino acid 116 to amino acid 125 of SEQ ID NO:210.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:209.

35 A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:

(i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(aa) SEQ ID NO:209, but excluding the poly(A) tail at the 3' end of SEQ ID NO:209; and

5 (ab) the nucleotide sequence of the cDNA insert of clone bh157\_7 deposited under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

(iii) isolating the DNA polynucleotide detected with the probe(s);  
and

10 (b) a process comprising the steps of:

(i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:209, but excluding the poly(A) tail at the 3' end of SEQ ID NO:209; and

15 (bb) the nucleotide sequence of the cDNA insert of clone bh157\_7 deposited under ATCC 98580;

(ii) hybridizing said primer(s) to human DNA;

(iii) amplifying human DNA sequences; and

(iv) isolating the polynucleotide product of step (b)(iii).

20 Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:209, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:209 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:209 but excluding the poly(A) tail at the 3' end of SEQ ID NO:209. In another preferred embodiment, the  
25 nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:209 from nucleotide 183 to nucleotide 911, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:209 from nucleotide 183 to nucleotide 911, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:209 from nucleotide 183 to nucleotide 911.

30 In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

(a) the amino acid sequence of SEQ ID NO:210;

(b) the amino acid sequence of SEQ ID NO:210 from amino acid 1 to amino  
35 acid 76;

(c) fragments of the amino acid sequence of SEQ ID NO:210 comprising eight consecutive amino acids of SEQ ID NO:210; and

(d) the amino acid sequence encoded by the cDNA insert of clone bh157\_7 deposited under accession number ATCC 98580; the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:210 or the amino acid  
 5 sequence of SEQ ID NO:210 from amino acid 1 to amino acid 76. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:210 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:210, or a protein comprising a fragment of the  
 10 amino acid sequence of SEQ ID NO:210 having biological activity, the fragment comprising the amino acid sequence from amino acid 116 to amino acid 125 of SEQ ID NO:210.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 15 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:211;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:211 from nucleotide 1391 to nucleotide 1609;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:211 from nucleotide 1439 to nucleotide 1609;
- 20 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone cg426\_8 deposited under accession number ATCC 98580;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone cg426\_8 deposited under accession number ATCC 98580;
- 25 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone cg426\_8 deposited under accession number ATCC 98580;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone cg426\_8 deposited under accession number ATCC 98580;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence  
 30 of SEQ ID NO:212;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:212 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:212;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g)  
 35 above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:211 from nucleotide 1391 to nucleotide 1609; the nucleotide sequence of SEQ ID NO:211 from nucleotide 1439 to nucleotide 1609; the nucleotide sequence of the full-length protein coding sequence of clone cg426\_8 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone cg426\_8 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone cg426\_8 deposited under accession number ATCC 98580. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:212 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:212, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:212 having biological activity, the fragment comprising the amino acid sequence from amino acid 31 to amino acid 40 of SEQ ID NO:212.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:211.

A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:

(i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(aa) SEQ ID NO:211, but excluding the poly(A) tail at the 3' end of SEQ ID NO:211; and

(ab) the nucleotide sequence of the cDNA insert of clone cg426\_8 deposited under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

(iii) isolating the DNA polynucleotide detected with the probe(s); and

(b) a process comprising the steps of:

(i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:211, but excluding the poly(A) tail at the 3' end of SEQ ID NO:211; and

(bb) the nucleotide sequence of the cDNA insert of clone cg426\_8 deposited under ATCC 98580;

- Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:211, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:211 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:211 but excluding the poly(A) tail at the 3' end of SEQ ID NO:211. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:211 from nucleotide 1391 to nucleotide 1609, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:211 from nucleotide 1391 to nucleotide 1609, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:211 from nucleotide 1391 to nucleotide 1609. In a further preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:211 from nucleotide 1439 to nucleotide 1609, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:211 from nucleotide 1439 to nucleotide 1609, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:211 from nucleotide 1439 to nucleotide 1609.

(a) the amino acid sequence of SEQ ID NO:212;

(b) fragments of the amino acid sequence of SEQ ID NO:212 comprising  
25 eight consecutive amino acids of SEQ ID NO:212; and

(c) the amino acid sequence encoded by the cDNA insert of clone cg426\_8  
deposited under accession number ATCC 98580;

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:



- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:213;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:213 from nucleotide 185 to nucleotide 586;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:213 from nucleotide 578 to nucleotide 586;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ck48\_12 deposited under accession number ATCC 98580;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ck48\_12 deposited under accession number ATCC 98580;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ck48\_12 deposited under accession number ATCC 98580;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ck48\_12 deposited under accession number ATCC 98580;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:214;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:214 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:214;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
- (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
- (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:213 from nucleotide 185 to nucleotide 586; the nucleotide sequence of SEQ ID NO:213 from nucleotide 578 to nucleotide 586; the nucleotide sequence of the full-length protein coding sequence of clone ck48\_12 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone ck48\_12 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ck48\_12 deposited under accession number ATCC 98580. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:214 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:214, or a polynucleotide encoding a protein comprising a fragment of the amino acid

sequence of SEQ ID NO:214 having biological activity, the fragment comprising the amino acid sequence from amino acid 62 to amino acid 71 of SEQ ID NO:214.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:213.

- 5 A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:
- (a) a process comprising the steps of:
    - (i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
      - 10 (aa) SEQ ID NO:213, but excluding the poly(A) tail at the 3' end of SEQ ID NO:213; and
      - (ab) the nucleotide sequence of the cDNA insert of clone ck48\_12 deposited under ATCC 98580;
      - (ii) hybridizing said probe(s) to human DNA; and
      - 15 (iii) isolating the DNA polynucleotide detected with the probe(s); and
    - (b) a process comprising the steps of:
      - (i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
        - 20 (ba) SEQ ID NO:213, but excluding the poly(A) tail at the 3' end of SEQ ID NO:213; and
        - (bb) the nucleotide sequence of the cDNA insert of clone ck48\_12 deposited under ATCC 98580;
        - (ii) hybridizing said primer(s) to human DNA;
        - 25 (iii) amplifying human DNA sequences; and
        - (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:213, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:213 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:9 but excluding the poly(A) tail at the 3' end of SEQ ID NO:213. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:213 from nucleotide 185 to nucleotide 586, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:213 from nucleotide 185 to nucleotide 586, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:213 from nucleotide 185 to nucleotide 586.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:214;
  - 5 (b) fragments of the amino acid sequence of SEQ ID NO:214 comprising eight consecutive amino acids of SEQ ID NO:214; and
  - (c) the amino acid sequence encoded by the cDNA insert of clone ck48\_12 deposited under accession number ATCC 98580;
- the protein being substantially free from other mammalian proteins. Preferably such
- 10 protein comprises the amino acid sequence of SEQ ID NO:214. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:214 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:214, or a protein comprising a fragment of the
  - 15 amino acid sequence of SEQ ID NO:214 having biological activity, the fragment comprising the amino acid sequence from amino acid 62 to amino acid 71 of SEQ ID NO:214.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- 20 (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:215;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:215 from nucleotide 554 to nucleotide 1012;
- (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:215 from nucleotide 632 to nucleotide 1012;
- 25 (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone co1000\_1 deposited under accession number ATCC 98580;
- (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone co1000\_1 deposited under accession number ATCC 98580;
- 30 (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone co1000\_1 deposited under accession number ATCC 98580;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone co1000\_1 deposited under accession number ATCC 98580;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence
- 35 of SEQ ID NO:216;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:216 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:216;

(j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

5 (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:215 from nucleotide 554 to nucleotide 1012; the nucleotide sequence of SEQ ID NO:215 from nucleotide 632 to nucleotide 1012; the nucleotide sequence of the full-length protein coding sequence of clone co1000\_1 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone co1000\_1 deposited under accession number ATCC 98580. In other preferred  
10 embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone co1000\_1 deposited under accession number ATCC 98580. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:216 from amino acid 1 to amino acid 63. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:216 having biological activity, the fragment preferably  
15 comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:216, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:216 having biological activity, the fragment comprising the amino acid sequence from amino acid 71 to amino acid 80 of SEQ ID NO:216.

25 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:215.

A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:  
30 (i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(aa) SEQ ID NO:215, but excluding the poly(A) tail at the 3' end of SEQ ID NO:215; and

(ab) the nucleotide sequence of the cDNA insert of clone co1000\_1 deposited  
35 under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

(iii) isolating the DNA polynucleotide detected with the probe(s);  
and

(b) a process comprising the steps of:

(i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:215, but excluding the poly(A) tail at the 3' end of SEQ ID  
5 NO:215; and

(bb) the nucleotide sequence of the cDNA insert of clone co1000\_1 deposited under ATCC 98580;

(ii) hybridizing said primer(s) to human DNA;

(iii) amplifying human DNA sequences; and

10 (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:215, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:215 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:215 but excluding  
15 the poly(A) tail at the 3' end of SEQ ID NO:215. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:215 from nucleotide 554 to nucleotide 1012, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:215 from nucleotide 554 to nucleotide 1012, to a nucleotide sequence corresponding  
20 to the 3' end of said sequence of SEQ ID NO:215 from nucleotide 554 to nucleotide 1012. In a further preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:215 from nucleotide 632 to nucleotide 1012, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:215 from nucleotide 632 to  
25 nucleotide 1012, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:215 from nucleotide 632 to nucleotide 1012.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

30 (a) the amino acid sequence of SEQ ID NO:216;

(b) the amino acid sequence of SEQ ID NO:216 from amino acid 1 to amino acid 63;

(c) fragments of the amino acid sequence of SEQ ID NO:216 comprising eight consecutive amino acids of SEQ ID NO:216; and

35 (d) the amino acid sequence encoded by the cDNA insert of clone co1000\_1 deposited under accession number ATCC 98580;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:216 or the amino acid

sequence of SEQ ID NO:216 from amino acid 1 to amino acid 63. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:216 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:216, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:216 having biological activity, the fragment comprising the amino acid sequence from amino acid 71 to amino acid 80 of SEQ ID NO:216.

- In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:
- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:217;
  - (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:217 from nucleotide 83 to nucleotide 1111;
  - (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:217 from nucleotide 155 to nucleotide 1111;
  - (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone ct489\_14 deposited under accession number ATCC 98580;
  - (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone ct489\_14 deposited under accession number ATCC 98580;
  - (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone ct489\_14 deposited under accession number ATCC 98580;
  - (g) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone ct489\_14 deposited under accession number ATCC 98580;
  - (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:218;
  - (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:218 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:218;
  - (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;
  - (k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and
  - (l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:217 from nucleotide 83 to nucleotide 1111; the nucleotide sequence of SEQ ID NO:217 from nucleotide 155 to nucleotide 1111; the nucleotide sequence of the full-

length protein coding sequence of clone ct489\_14 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone ct489\_14 deposited under accession number ATCC 98580. In other preferred  
 5 embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone ct489\_14 deposited under accession number ATCC 98580. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:218 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:218,  
 10 or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:218 having biological activity, the fragment comprising the amino acid sequence from amino acid 166 to amino acid 175 of SEQ ID NO:218.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:217.

- 15 A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:
- (a) a process comprising the steps of:
    - (i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
      - 20 (aa) SEQ ID NO:217, but excluding the poly(A) tail at the 3' end of SEQ ID NO:217; and
      - (ab) the nucleotide sequence of the cDNA insert of clone ct489\_14 deposited under ATCC 98580;
    - (ii) hybridizing said probe(s) to human DNA; and
    - 25 (iii) isolating the DNA polynucleotide detected with the probe(s); and
  - (b) a process comprising the steps of:
    - (i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
      - 30 (ba) SEQ ID NO:217, but excluding the poly(A) tail at the 3' end of SEQ ID NO:217; and
      - (bb) the nucleotide sequence of the cDNA insert of clone ct489\_14 deposited under ATCC 98580;
    - (ii) hybridizing said primer(s) to human DNA;
    - 35 (iii) amplifying human DNA sequences; and
    - (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:217, and extends

contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:217 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:217 but excluding the poly(A) tail at the 3' end of SEQ ID NO:217. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:217 from nucleotide 83 to nucleotide 1111, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:217 from nucleotide 83 to nucleotide 1111, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:217 from nucleotide 83 to nucleotide 1111. In a further preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:217 from nucleotide 155 to nucleotide 1111, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:217 from nucleotide 155 to nucleotide 1111, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:217 from nucleotide 155 to nucleotide 1111.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:218;
- (b) fragments of the amino acid sequence of SEQ ID NO:218 comprising eight consecutive amino acids of SEQ ID NO:218; and
- (c) the amino acid sequence encoded by the cDNA insert of clone ct489\_14 deposited under accession number ATCC 98580;

the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:218. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:218 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:218, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:218 having biological activity, the fragment comprising the amino acid sequence from amino acid 166 to amino acid 175 of SEQ ID NO:218.

In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:219;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:219 from nucleotide 26 to nucleotide 490;



(c) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone df821\_1 deposited under accession number ATCC 98580;

(d) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone df821\_1 deposited under accession number ATCC 98580;

(e) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone df821\_1 deposited under accession number ATCC 98580;

(f) a polynucleotide encoding a mature protein encoded by the cDNA insert of clone df821\_1 deposited under accession number ATCC 98580;

(g) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:220;

(h) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:220 having biological activity, the fragment comprising eight consecutive amino acids of SEQ ID NO:220;

(i) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(f) above;

(j) a polynucleotide which encodes a species homologue of the protein of (g) or (h) above; and

(k) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(h).

Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:219 from nucleotide 26 to nucleotide 490; the nucleotide sequence of the full-length protein coding sequence of clone df821\_1 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone df821\_1 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone df821\_1 deposited under accession number ATCC 98580. In yet other preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:220 from amino acid 92 to amino acid 152. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:220 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:220, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:220 having biological activity, the fragment comprising the amino acid sequence from amino acid 72 to amino acid 81 of SEQ ID NO:220.

Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:219.

A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

- (a) a process comprising the steps of:
  - (i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
    - (aa) SEQ ID NO:219, but excluding the poly(A) tail at the 3' end of SEQ ID NO:219; and
    - (ab) the nucleotide sequence of the cDNA insert of clone df821\_1 deposited under ATCC 98580;
  - (ii) hybridizing said probe(s) to human DNA; and
  - (iii) isolating the DNA polynucleotide detected with the probe(s); and
- (b) a process comprising the steps of:
  - (i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:
    - (ba) SEQ ID NO:219, but excluding the poly(A) tail at the 3' end of SEQ ID NO:15; and
    - (bb) the nucleotide sequence of the cDNA insert of clone df821\_1 deposited under ATCC 98580;
  - (ii) hybridizing said primer(s) to human DNA;
  - (iii) amplifying human DNA sequences; and
  - (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:219, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:219 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:219 but excluding the poly(A) tail at the 3' end of SEQ ID NO:219. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:219 from nucleotide 26 to nucleotide 490, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:219 from nucleotide 26 to nucleotide 490, to a nucleotide sequence corresponding to the 3' end of said sequence of SEQ ID NO:219 from nucleotide 26 to nucleotide 490.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:220;
- (b) the amino acid sequence of SEQ ID NO:220 from amino acid 92 to amino acid 152;

(c) fragments of the amino acid sequence of SEQ ID NO:220 comprising eight consecutive amino acids of SEQ ID NO:220; and

(d) the amino acid sequence encoded by the cDNA insert of clone df821\_1 deposited under accession number ATCC 98580;

- 5 the protein being substantially free from other mammalian proteins. Preferably such protein comprises the amino acid sequence of SEQ ID NO:220 or the amino acid sequence of SEQ ID NO:220 from amino acid 92 to amino acid 152. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:220 having biological activity, the fragment
- 10 preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:220, or a protein comprising a fragment of the amino acid sequence of SEQ ID NO:220 having biological activity, the fragment comprising the amino acid sequence from amino acid 72 to amino acid 81 of SEQ ID NO:220.

- 15 In one embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of:

- (a) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:221;
- (b) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:221 from nucleotide 65 to nucleotide 412;
- 20 (c) a polynucleotide comprising the nucleotide sequence of SEQ ID NO:221 from nucleotide 197 to nucleotide 412;
- (d) a polynucleotide comprising the nucleotide sequence of the full-length protein coding sequence of clone dy41\_2 deposited under accession number ATCC 98580;
- 25 (e) a polynucleotide encoding the full-length protein encoded by the cDNA insert of clone dy41\_2 deposited under accession number ATCC 98580;
- (f) a polynucleotide comprising the nucleotide sequence of a mature protein coding sequence of clone dy41\_2 deposited under accession number ATCC 98580;
- (g) a polynucleotide encoding a mature protein encoded by the cDNA insert
- 30 of clone dy41\_2 deposited under accession number ATCC 98580;
- (h) a polynucleotide encoding a protein comprising the amino acid sequence of SEQ ID NO:222;
- (i) a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:222 having biological activity, the fragment comprising
- 35 eight consecutive amino acids of SEQ ID NO:222;
- (j) a polynucleotide which is an allelic variant of a polynucleotide of (a)-(g) above;

(k) a polynucleotide which encodes a species homologue of the protein of (h) or (i) above ; and

(l) a polynucleotide that hybridizes under stringent conditions to any one of the polynucleotides specified in (a)-(i).

5 Preferably, such polynucleotide comprises the nucleotide sequence of SEQ ID NO:221 from nucleotide 65 to nucleotide 412; the nucleotide sequence of SEQ ID NO:221 from nucleotide 197 to nucleotide 412; the nucleotide sequence of the full-length protein coding sequence of clone dy41\_2 deposited under accession number ATCC 98580; or the nucleotide sequence of a mature protein coding sequence of clone  
10 dy41\_2 deposited under accession number ATCC 98580. In other preferred embodiments, the polynucleotide encodes the full-length or a mature protein encoded by the cDNA insert of clone dy41\_2 deposited under accession number ATCC 98580. In further preferred embodiments, the present invention provides a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID  
15 NO:222 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:222, or a polynucleotide encoding a protein comprising a fragment of the amino acid sequence of SEQ ID NO:222 having biological activity, the fragment comprising the amino acid sequence from amino acid 53 to amino acid 62 of SEQ ID NO:222.

20 Other embodiments provide the gene corresponding to the cDNA sequence of SEQ ID NO:221.

A further embodiment of the invention provides a process for producing an isolated polynucleotide, wherein the process is selected from the group consisting of:

(a) a process comprising the steps of:

25 (i) preparing one or more polynucleotide probes that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(aa) SEQ ID NO:221, but excluding the poly(A) tail at the 3' end of SEQ ID NO:221; and

30 (ab) the nucleotide sequence of the cDNA insert of clone dy41\_2 deposited under ATCC 98580;

(ii) hybridizing said probe(s) to human DNA; and

(iii) isolating the DNA polynucleotide detected with the probe(s);  
and

(b) a process comprising the steps of:

35 (i) preparing one or more polynucleotide primers that hybridize in 6X SSC at 65 degrees C to a nucleotide sequence selected from the group consisting of:

(ba) SEQ ID NO:221, but excluding the poly(A) tail at the 3' end of SEQ ID NO:221; and

(bb) the nucleotide sequence of the cDNA insert of clone dy41\_2 deposited under ATCC 98580;

(ii) hybridizing said primer(s) to human DNA;

(iii) amplifying human DNA sequences; and

5 (iv) isolating the polynucleotide product of step (b)(iii).

Preferably, the nucleotide sequence of the polynucleotide isolated according to this method corresponds to the cDNA sequence of SEQ ID NO:221, and extends contiguously from a nucleotide sequence corresponding to the 5' end of SEQ ID NO:221 to a nucleotide sequence corresponding to the 3' end of SEQ ID NO:221 but excluding  
10 the poly(A) tail at the 3' end of SEQ ID NO:221. In another preferred embodiment, the nucleotide sequence of said isolated polynucleotide corresponds to the cDNA sequence of SEQ ID NO:221 from nucleotide 65 to nucleotide 412, and extends contiguously from a nucleotide sequence corresponding to the 5' end of said sequence of SEQ ID NO:221 from nucleotide 65 to nucleotide 412, to a nucleotide sequence corresponding to the 3'  
15 end of said sequence of SEQ ID NO:221 from nucleotide 65 to nucleotide 412.

In other embodiments, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence of SEQ ID NO:222;
  - 20 (b) fragments of the amino acid sequence of SEQ ID NO:222 comprising eight consecutive amino acids of SEQ ID NO:222; and
  - (c) the amino acid sequence encoded by the cDNA insert of clone dy41\_2 deposited under accession number ATCC 98580;
- the protein being substantially free from other mammalian proteins. Preferably such  
25 protein comprises the amino acid sequence of SEQ ID NO:222. In further preferred embodiments, the present invention provides a protein comprising a fragment of the amino acid sequence of SEQ ID NO:222 having biological activity, the fragment preferably comprising eight (more preferably twenty, most preferably thirty) consecutive amino acids of SEQ ID NO:222, or a protein comprising a fragment of the  
30 amino acid sequence of SEQ ID NO:222 having biological activity, the fragment comprising the amino acid sequence from amino acid 53 to amino acid 62 of SEQ ID NO:222.

In certain preferred embodiments, the polynucleotide is operably linked to an expression control sequence. The invention also provides a host cell, including  
35 bacterial, yeast, insect and mammalian cells, transformed with such polynucleotide compositions. Also provided by the present invention are organisms that have enhanced, reduced, or modified expression of the gene(s) corresponding to the polynucleotide sequences disclosed herein.

Processes are also provided for producing a protein encoded by polynucleotides of the present invention, which comprise:

(a) growing a culture of the host cell transformed with such polynucleotides in a suitable culture medium; and

5 (b) purifying the protein from the culture.

The protein produced according to such methods is also provided by the present invention, as are isolated polynucleotides encoding the proteins produced according to such methods.

Protein compositions of the present invention may further comprise a  
10 pharmaceutically acceptable carrier. Compositions comprising an antibody which specifically reacts with such protein are also provided by the present invention.

Methods are also provided for preventing, treating or ameliorating a medical condition which comprises administering to a mammalian subject a therapeutically effective amount of a composition comprising a protein of the present invention and a  
15 pharmaceutically acceptable carrier.

#### BRIEF DESCRIPTION OF THE DRAWINGS

20 Figures 1A and 1B are schematic representations of the pED6 and pNOT's vectors, respectively, used for deposit of clones disclosed herein.

#### DETAILED DESCRIPTION

#### 25 ISOLATED PROTEINS AND POLYNUCLEOTIDES

Nucleotide and amino acid sequences, as presently determined, are reported below for each clone and protein disclosed in the present application. The nucleotide sequence of each clone can readily be determined by sequencing of the deposited clone in accordance with known methods. The predicted amino acid sequence (both full-  
30 length and mature forms) can then be determined from such nucleotide sequence. The amino acid sequence of the protein encoded by a particular clone can also be determined by expression of the clone in a suitable host cell, collecting the protein and determining its sequence. For each disclosed protein applicants have identified what they have determined to be the reading frame best identifiable with sequence  
35 information available at the time of filing.

As used herein a "secreted" protein is one which, when expressed in a suitable host cell, is transported across or through a membrane, including transport as a result of signal sequences in its amino acid sequence. "Secreted" proteins include without

limitation proteins secreted wholly (e.g., soluble proteins) or partially (e.g., receptors) from the cell in which they are expressed. "Secreted" proteins also include without limitation proteins which are transported across the membrane of the endoplasmic reticulum.

5

Clone "bh389\_11"

A polynucleotide of the present invention has been identified as clone "bh389\_11". bh389\_11 was isolated from a human adult thyroid cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bh389\_11 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bh389\_11 protein").

The nucleotide sequence of bh389\_11 as presently determined is reported in SEQ ID NO:1. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bh389\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:2. Amino acids 10 to 22 of SEQ ID NO:2 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 23, or are a transmembrane domain. The TopPredII computer program predicts a potential transmembrane domain within the bh389\_11 protein sequence centered around amino acid 68 of SEQ ID NO:2.

Another potential bh389\_11 reading frame and predicted amino acid sequence is encoded by basepairs 757 to 1833 of SEQ ID NO:1 and is reported in SEQ ID NO:34. A frameshift in the nucleotide sequence of SEQ ID NO:1 between about nucleotide 754 to about nucleotide 803 could join the reading frames of SEQ ID NO:1 and SEQ ID NO:34. The TopPredII computer program predicts a potential transmembrane domain within the amino acid sequence of SEQ ID NO:34, centered around amino acid 357 of SEQ ID NO:34.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bh389\_11 should be approximately 1700 bp.

The nucleotide sequence disclosed herein for bh389\_11 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bh389\_11 demonstrated at least some similarity with sequences identified as AA307880 (EST178733 Colon carcinoma (HCC) cell line Homo sapiens cDNA 5' end), AA442426 (zv70f06.r1 Soares total fetus Nb2HF8 9w Homo sapiens cDNA clone 759011 5'), H70103 (yr92f04.r1 Homo sapiens cDNA clone 212767 5'), R19820 (yg37f12.r1 Homo sapiens cDNA clone 34771 5'), and W46238 (zc30e10.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA clone 323850 3').

Based upon sequence similarity, bh389\_11 proteins and each similar protein or peptide may share at least some activity.

#### Clone "bk112\_15"

5 A polynucleotide of the present invention has been identified as clone "bk112\_15". bk112\_15 was isolated from a human adult retina cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein.  
10 bk112\_15 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bk112\_15 protein").

The nucleotide sequence of bk112\_15 as presently determined is reported in SEQ ID NO:3. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bk112\_15 protein corresponding to the  
15 foregoing nucleotide sequence is reported in SEQ ID NO:4.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bk112\_15 should be approximately 1300 bp.

The nucleotide sequence disclosed herein for bk112\_15 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
20 FASTA search protocols. bk112\_15 demonstrated at least some similarity with sequences identified as AA307119 (EST178031 Colon carcinoma (HCC) cell line Homo sapiens cDNA 5' end), AA318352 (EST20422 Retina II Homo sapiens cDNA 5' end similar to similar to C. elegans hypothetical protein, cosmid ZK688.2), L20941 (Human ferritin heavy chain mRNA, complete cds), M97164 (Human ferritin heavy  
25 chain mRNA, complete cds), N25339 (yx55d08.s1 Homo sapiens cDNA clone 265647 3'), N31453 (yx55d08.r1 Homo sapiens cDNA clone 265647 5'), and N33227 (yy07d02.s1 Homo sapiens cDNA clone 270531 3' similar to gb:L20941 FERRITIN HEAVY CHAIN (HUMAN)). The predicted amino acid sequence disclosed herein for bk112\_15 was searched against the GenPept and GeneSeq amino acid sequence  
30 databases using the BLASTX search protocol. The predicted bk112\_15 protein demonstrated at least some similarity to sequences identified as Z68335 (C29F4.2 [Caenorhabditis elegans]). Based upon sequence similarity, bk112\_15 proteins and each similar protein or peptide may share at least some activity.

#### Clone "bk200\_13"

35 A polynucleotide of the present invention has been identified as clone "bk200\_13". bk200\_13 was isolated from a human adult retina cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No.



5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bk200\_13 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bk200\_13 protein").

5 The nucleotide sequence of bk200\_13 as presently determined is reported in SEQ ID NO:5. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bk200\_13 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:6.

10 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bk200\_13 should be approximately 1000 bp.

The nucleotide sequence disclosed herein for bk200\_13 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bk200\_13 demonstrated at least some similarity with sequences identified as AA098915 zk84f06.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 489539 3'), AA150367 zl07b06.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 491603 5'), AA235904 (zs40h05.r1 Soares NhHMPu S1 Homo sapiens cDNA clone 687705 5'), N32487 (yx79g10.r1 Homo sapiens cDNA clone 268002 5'), and T47862 (yb17g03.r1 Homo sapiens cDNA clone 71476 5'). Based upon sequence similarity, bk200\_13 proteins and each similar protein or peptide may share at least some activity. The nucleotide sequence of bk200\_13 may contain CAAAAA repeat-like elements.

#### Clone "di386\_3"

25 A polynucleotide of the present invention has been identified as clone "di386\_3". di386\_3 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. di386\_3 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "di386\_3 protein").

30 The nucleotide sequence of the 5' portion of di386\_3 as presently determined is reported in SEQ ID NO:7. What applicants presently believe is the proper reading frame for the coding region is indicated in SEQ ID NO:8. The predicted amino acid sequence of the di386\_3 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:8. Amino acids 39 to 51 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 52, or are a transmembrane domain. Amino acids 17 to 29 OF SEQ ID NO:8 are also a possible leader/signal sequence, with the predicted mature amino acid sequence

beginning at amino acid 30, or are a transmembrane domain. Additional nucleotide sequence from the 3' portion of di386\_3, including the polyA tail, is reported in SEQ ID NO:9.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone di386\_3 should be approximately 2000 bp.

The nucleotide sequence disclosed herein for di386\_3 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. di386\_3 demonstrated no similarity with any known sequences in those databases.

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#### Clone "em397\_2"

A polynucleotide of the present invention has been identified as clone "em397\_2". em397\_2 was isolated from a human fetal kidney cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. em397\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "em397\_2 protein").

The nucleotide sequence of em397\_2 as presently determined is reported in SEQ ID NO:10. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the em397\_2 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:11.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone em397\_2 should be approximately 1250 bp.

The nucleotide sequence disclosed herein for em397\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. em397\_2 demonstrated at least some similarity with sequences identified as AA092876 (m0851.seq.F Fetal heart, Lambda ZAP Express Homo sapiens cDNA 5'), AA180952 (zp41b06.r1 Stratagene muscle 937209 Homo sapiens cDNA clone 611987 5'), AA463323 (zx71f01.r1 Soares total fetus Nb2HF8 9w Homo sapiens), H87081 (ys74f01.r1 Homo sapiens cDNA clone 220537 5'), W56381 (zc57a01.r1 Soares parathyroid tumor NbHPA Homo sapiens cDNA clone 326376 5'), W88527 (zh73g02.s1 Soares fetal liver spleen 1NFLS S1 Homo sapiens cDNA clone 417746 3'), and Z64565 (H.sapiens CpG island DNA genomic MseI fragment, clone 13d12, reverse read cpg13d12.r1c). Based upon sequence similarity, em397\_2 proteins and each similar protein or peptide may share at least some activity.

Clone "fh170\_7"

A polynucleotide of the present invention has been identified as clone "fh170\_7". fh170\_7 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fh170\_7 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "fh170\_7 protein").

The nucleotide sequence of fh170\_7 as presently determined is reported in SEQ ID NO:12. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the fh170\_7 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:13. Amino acids 127 to 139 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 140, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fh170\_7 should be approximately 2200 bp.

The nucleotide sequence disclosed herein for fh170\_7 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fh170\_7 demonstrated at least some similarity with sequences identified as AA112479 (zn69a02.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone 563402 3'), AA593402 (nn57g10.s1 NCI\_CGAP\_Kid6 Homo sapiens cDNA clone IMAGE:1088034), Q76795 (Human genome fragment), T26136 (Human gene signature HUMGS08373), and Z19759 (H. sapiens putatively transcribed partial sequence; UK-HGMP sequence ID AAAALWX; single read). The predicted amino acid sequence disclosed herein for fh170\_7 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted fh170\_7 protein demonstrated at least some similarity to sequences identified as D32253 (MagA [Magnetospirillum sp.]) and W01520 (MagA protein). The predicted fh170\_7 protein also demonstrated at least some similarity to other prokaryotic membrane transport proteins: potassium-efflux system protein kefB and NaH-antiporter protein. Based upon sequence similarity, fh170\_7 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts ten potential transmembrane domains within the fh170\_7 protein sequence, centered around amino acids 130, 160, 210, 230, 280, 310, 360, 380, 420, and 500 of SEQ ID NO:13, respectively.

Clone "fn53\_4"

A polynucleotide of the present invention has been identified as clone "fn53\_4". fn53\_4 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fn53\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "fn53\_4 protein").

The nucleotide sequence of the 5' portion of fn53\_4 as presently determined is reported in SEQ ID NO:14. An additional internal nucleotide sequence from fn53\_4 as presently determined is reported in SEQ ID NO:15. What applicants believe is the proper reading frame and the predicted amino acid sequence encoded by such internal sequence is reported in SEQ ID NO:16. Additional nucleotide sequence from the 3' portion of fn53\_4, including the polyA tail, is reported in SEQ ID NO:17.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fn53\_4 should be approximately 4100 bp.

The nucleotide sequence disclosed herein for fn53\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fn53\_4 demonstrated at least some similarity with sequences identified as AA179207 (zp46c11.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone 612500 3'), AA279207 (zs83e06.s1 NCI\_CGAP\_GCB1 Homo sapiens cDNA clone IMAGE:704098 3', mRNA sequence), H87151 (yw15a06.s1 Homo sapiens cDNA clone 252274 3'), and H83373 (ys90a09.r1 Homo sapiens cDNA clone 222040 5' similar to SP:BICD\_DROME P16568 CYTOSKELETON-LIKE BICAUDAL D). The predicted amino acid sequence disclosed herein for fn53\_4 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted fn53\_4 protein demonstrated at least some similarity to sequences identified as M31684 and X51652 (bicaudalD protein [Drosophila melanogaster]) and R66930 (AMML chromosome inv(16) product). Based upon sequence similarity, fn53\_4 proteins and each similar protein or peptide may share at least some activity.

Clone "fq505\_4"

A polynucleotide of the present invention has been identified as clone "fq505\_4". fq505\_4 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fq505\_4 is a full-length

clone, including the entire coding sequence of a secreted protein (also referred to herein as "fq505\_4 protein").

The nucleotide sequence of fq505\_4 as presently determined is reported in SEQ ID NO:18. What applicants presently believe to be the proper reading frame and the  
5 predicted amino acid sequence of the fq505\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:19.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fq505\_4 should be approximately 512 bp.

The nucleotide sequence disclosed herein for fq505\_4 was searched against the  
10 GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fq505\_4 demonstrated at least some similarity with sequences identified as Z71861 (C.hircus mRNA for EST2-31). The predicted amino acid sequence disclosed herein for fq505\_4 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted  
15 fq505\_4 protein demonstrated at least some similarity to sequences identified as P92141 (Recombinant human adult T cell leukaemia derived factor polypeptide), X54539 (thioredoxin [Homo sapiens]), and X77584 (ATL-derived factor/thioredoxin [Homo sapiens]). The predicted fq505\_4 protein also demonstrated at least some similarity to sequences identified as surface associated sulphydryl protein (GenProt accession  
20 number135773). The similarity between these proteins includes a WCGPC catalytic site, which is present as RCGPC at amino acids 31 to 35 of the predicted fq505\_4 protein. In addition to having thioredoxin catalytic activity, at least one thioredoxin-related protein has also been reported to be "an IL-2 receptor/Tac inducer" (Tagaya *et al.*, 1989, *EMBO J.* **8**(3): 757-764). At least one thioredoxin-related protein is reported to  
25 be associated with the plasma membrane, "indicating that the protein may be a member of this [thioredoxin] family and function as an essential growth factor" (Martin and Dean, 1991, *Biochem. Biophys. Res. Commun.* **175**(1): 123-128). Based upon sequence similarity, fq505\_4 proteins and each similar protein or peptide may share at least some activity.

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#### Clone "fw13\_9"

A polynucleotide of the present invention has been identified as clone "fw13\_9". fw13\_9 was isolated from a human adult testes (teratocarcinoma NCCIT) cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat.  
35 No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fw13\_9 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "fw13\_9 protein").

The nucleotide sequence of fw13\_9 as presently determined is reported in SEQ ID NO:20. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the fw13\_9 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:21.

5           The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fw13\_9 should be approximately 1900 bp.

          The nucleotide sequence disclosed herein for fw13\_9 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fw13\_9 demonstrated at least some similarity with sequences  
10 identified as AA047557 (zf13f08.r1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 376839 5'), AA284524 (zt20d07.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 713677 3'), AA502778 (ne43e04.s1 NCI\_CGAP\_Co3 Homo sapiens cDNA clone IMAGE:900126), J04743 (M.musculus Ms6-hm locus, repeat elements), R35040 (yh86a10.r1 Homo sapiens cDNA clone 136602 5'), T21414 (Human gene  
15 signature HUMGS02783), and U91318 (Human chromosome 16p13 BAC clone CIT987SK-962B4 complete sequence). Based upon sequence similarity, fw13\_9 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the fw13\_9 protein sequence centered around amino acid 30 of SEQ ID NO:21.

#### 20           Clone "gg619\_2"

          A polynucleotide of the present invention has been identified as clone "gg619\_2". gg619\_2 was isolated from a human fetal kidney cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No.  
25 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. gg619\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "gg619\_2 protein").

          The nucleotide sequence of gg619\_2 as presently determined is reported in SEQ  
30 ID NO:22. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the gg619\_2 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:23.

          The EcoRI/NotI restriction fragment obtainable from the deposit containing clone gg619\_2 should be approximately 1350 bp.

35           The nucleotide sequence disclosed herein for gg619\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. gg619\_2 demonstrated at least some similarity with sequences identified as N42957 (yy12b12.r1 Homo sapiens cDNA clone 271007 5'

similar to SW:ALG5\_YEAST P40350 dolichyl-phosphate beta-glucosyltransferase), N50844 (yy91g05.s1 Homo sapiens cDNA clone 280952 3' similar to SW:ALG5\_YEAST P40350 dolichyl-phosphate beta-glucosyltransferase), and N62597 (yz75a06.s1 Homo sapiens cDNA clone 288850 3' similar to SW:ALG5\_YEAST P40350 Dolichyl-phosphate beta-glucosyltransferase). The predicted amino acid sequence disclosed herein for gg619\_2 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted gg619\_2 protein demonstrated at least some similarity to sequences identified as R38093 (nodC N-terminal portion [Bradyrhizobium sp. (Parasponia)]) and X77573 (dolichyl-phosphate beta-glucosyl-transferase [Saccharomyces cerevisiae]). The enzyme UDP-glucose:dolichyl-phosphate glucosyltransferase is a transmembrane-bound enzyme of the endoplasmic reticulum involved in protein N-linked glycosylation, and catalyzes the transfer of glucose from UDP-glucose to dolichyl phosphate. Based upon sequence similarity, gg619\_2 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the gg619\_2 protein sequence, centered around amino acid 188 of SEQ ID NO:23.

#### Clone "cl181\_3"

A polynucleotide of the present invention has been identified as clone "cl181\_3". cl181\_3 was isolated from a human adult retina cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cl181\_3 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cl181\_3 protein").

The nucleotide sequence of cl181\_3 as presently determined is reported in SEQ ID NO:35. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cl181\_3 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:36. Amino acids 50 to 62 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 63, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cl181\_3 should be approximately 2800 bp.

The nucleotide sequence disclosed herein for cl181\_3 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cl181\_3 demonstrated at least some similarity with sequences identified as C16190 (Human aorta cDNA 5'-end GEN-241B04) and L36900

(*Saccharomyces cerevisiae* mitochondrion transfer RNA-Ser1 (tRNA-Ser) gene and varI gene, complete cds). Based upon sequence similarity, cl181\_3 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the cl181\_3 protein sequence  
 5 centered around amino acid 77 of SEQ ID NO:36.

Clone "cr1044\_1"

A polynucleotide of the present invention has been identified as clone "cr1044\_1". cr1044\_1 was isolated from a human adult testes cDNA library using  
 10 methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cr1044\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cr1044\_1 protein").

15 The nucleotide sequence of cr1044\_1 as presently determined is reported in SEQ ID NO:37. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cr1044\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:38. Amino acids 158 to 170 of SEQ ID NO:36 are a predicted leader/signal sequence, with the predicted mature amino acid  
 20 sequence beginning at amino acid 171 of SEQ ID NO:38, or are a transmembrane domain. Base pairs 175 to 237 of SEQ ID NO:37 are a possible intron; if this sequence were removed from SEQ ID NO:37, another potential cr1044\_1 reading frame and predicted amino acid sequence that could be encoded by basepairs 45 to 830 of SEQ ID NO:37 is reported in SEQ ID NO:64. Amino acids 7 to 19 of SEQ ID NO:31 are a  
 25 predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 20 of SEQ ID NO:64, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cr1044\_1 should be approximately 3200 bp.

30 The nucleotide sequence disclosed herein for cr1044\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cr1044\_1 demonstrated at least some similarity with sequences identified as N99156 (zb81g04.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA clone 310038 3'), Q46852 (clone of recombinant human kappa casein gene fragment), and T20727 (Human gene signature HUMGS01945). The predicted  
 35 amino acid sequence disclosed herein for cr1044\_1 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted cr1044\_1 protein demonstrated at least some similarity to sequences identified as M16279 (antigen [Homo sapiens]) and U82164 (human CD99 type II).



Based upon sequence similarity, cr1044\_1 proteins and each similar protein or peptide may share at least some activity.

Clone "cz251\_1"

5 A polynucleotide of the present invention has been identified as clone "cz251\_1". cz251\_1 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cz251\_1 is a full-length  
10 clone, including the entire coding sequence of a secreted protein (also referred to herein as "cz251\_1 protein").

The nucleotide sequence of cz251\_1 as presently determined is reported in SEQ ID NO:39. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cz251\_1 protein corresponding to the foregoing  
15 nucleotide sequence is reported in SEQ ID NO:40.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cz251\_1 should be approximately 1200 bp.

The nucleotide sequence disclosed herein for cz251\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
20 FASTA search protocols. cz251\_1 demonstrated at least some similarity with sequences identified as R55084 (yg87a06.r1 Homo sapiens cDNA clone 40244 5') and U00930 (Human clone C4E 1.63 (CAC)<sub>n</sub>/(GTG)<sub>n</sub> repeat-containing mRNA). The predicted amino acid sequence disclosed herein for cz251\_1 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The  
25 predicted cz251\_1 protein demonstrated at least some similarity to the sequence identified as Z68751 (F01G4.1 [Caenorhabditis elegans]). Based upon sequence similarity, cz251\_1 proteins and each similar protein or peptide may share at least some activity. The nucleotide sequence of cz251\_1 may contain CAAA-like repeats.

Clone "dd12\_7"

30 A polynucleotide of the present invention has been identified as clone "dd12\_7". dd12\_7 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer  
35 analysis of the amino acid sequence of the encoded protein. dd12\_7 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "dd12\_7 protein").

The nucleotide sequence of dd12\_7 as presently determined is reported in SEQ ID NO:41. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the dd12\_7 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:42.

- 5 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone dd12\_7 should be approximately 1550 bp.

The nucleotide sequence disclosed herein for dd12\_7 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. dd12\_7 demonstrated at least some similarity with sequences identified as AA257999 (zs34a02.s1 Soares NbHTGBC Homo sapiens cDNA clone 10 687050 3'). Based upon sequence similarity, dd12\_7 proteins and each similar protein or peptide may share at least some activity.

#### Clone "fn191\_3"

- 15 A polynucleotide of the present invention has been identified as clone "fn191\_3". fn191\_3 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fn191\_3 is a full-length 20 clone, including the entire coding sequence of a secreted protein (also referred to herein as "fn191\_3 protein").

The nucleotide sequence of fn191\_3 as presently determined is reported in SEQ ID NO:43. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the fn191\_3 protein corresponding to the foregoing 25 nucleotide sequence is reported in SEQ ID NO:44. Amino acids 39 to 51 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 52, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fn191\_3 should be approximately 3000 bp.

- 30 The nucleotide sequence disclosed herein for fn191\_3 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fn191\_3 demonstrated at least some similarity with sequences identified as AA046787 (zk72f07.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 488389 5' similar to gb:L07077 ENOYL-COA HYDRATASE 35 (HUMAN);contains Alu repetitive element), G13132 (human chromosome 7 STS sWSS3349; single read), T06013 (EST03902 Homo sapiens cDNA clone HFBDL25), T08594 (EST06486 Homo sapiens cDNA clone HIBBG72 5' end), and W28342 (45g9 Human retina cDNA randomly primed sublibrary Homo sapiens cDNA). Based upon

sequence similarity, fn191\_3 proteins and each similar protein or peptide may share at least some activity. The nucleotide sequence in the 5' untranslated region of fn191\_3 may contain some repetitive elements.

5           Clone "gm196\_4"

A polynucleotide of the present invention has been identified as clone "gm196\_4". gm196\_4 was isolated from a human adult uterus cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the  
10 basis of computer analysis of the amino acid sequence of the encoded protein. gm196\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "gm196\_4 protein").

The nucleotide sequence of gm196\_4 as presently determined is reported in SEQ ID NO:45. What applicants presently believe to be the proper reading frame and the  
15 predicted amino acid sequence of the gm196\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:46. Another potential gm196\_4 reading frame and predicted amino acid sequence is encoded by basepairs 1364 to 2809 of SEQ ID NO:43 and is reported in SEQ ID NO:66. A frameshift in the nucleotide sequence of SEQ ID NO:45 could join the reading frames of SEQ ID NO:46 and SEQ ID NO:66. The  
20 TopPredII computer program predicts two potential transmembrane domains within the amino acid sequence of SEQ ID NO:66. Preferred fragments of the amino acid sequence of SEQ ID NO:66 comprise amino acids 1 to 163 or amino acids 236 to 245 of SEQ ID NO:66.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
25 clone gm196\_4 should be approximately 3000 bp.

The nucleotide sequence disclosed herein for gm196\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. gm196\_4 demonstrated at least some similarity with sequences identified as AA233552 (zr43a10.s1 Soares NhHMPu S1 Homo sapiens  
30 cDNA clone 666138 3'), AB005666 (Homo sapiens mRNA for GTPase-activating protein, complete cds), F12887 (H. sapiens partial cDNA sequence; clone c-3fh09), T25078 (Human gene signature HUMGS07218), and T75264 (yc88g09.r1 Homo sapiens cDNA clone 23008 5'). The predicted amino acid sequence disclosed herein for gm196\_4 was searched against the GenPept and GeneSeq amino acid sequence  
35 databases using the BLASTX search protocol. The predicted gm196\_4 protein demonstrated at least some similarity to the sequence identified as M64788 (GTPase activating protein [Homo sapiens]). Based upon sequence similarity, gm196\_4 proteins and each similar protein or peptide may share at least some activity.

Clone "gn114\_1"

A polynucleotide of the present invention has been identified as clone "gn114\_1". gn114\_1 was isolated from a human adult blood (peripheral blood mononuclear cells treated with granulocyte-colony stimulating factor *in vivo*) cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. gn114\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "gn114\_1 protein").

The nucleotide sequence of gn114\_1 as presently determined is reported in SEQ ID NO:47. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the gn114\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:48. Amino acids 26 to 38 of SEQ ID NO:48 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 39 of SEQ ID NO:48, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone gn114\_1 should be approximately 1500 bp.

The nucleotide sequence disclosed herein for gn114\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. gn114\_1 demonstrated at least some similarity with sequences identified as AA370547 (EST82206 Prostate gland I Homo sapiens cDNA 5' end), C04732 (Human Heart cDNA, clone 3NHC3910), and C05361 (Human Heart cDNA, clone 3NHC2451). Based upon sequence similarity, gn114\_1 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts an additional potential transmembrane domain within the gn114\_1 protein sequence centered around amino acid 90 of SEQ ID NO:48.

Clone "hj968\_2"

A polynucleotide of the present invention has been identified as clone "hj968\_2". hj968\_2 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. hj968\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "hj968\_2 protein").

The nucleotide sequence of hj968\_2 as presently determined is reported in SEQ ID NO:49. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the hj968\_2 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:50. Amino acids 1 to 9 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 10, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone hj968\_2 should be approximately 1800 bp.

The nucleotide sequence disclosed herein for hj968\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. hj968\_2 demonstrated at least some similarity with sequences identified as AA071746 (mf16h07.r1 Life Tech mouse brain Mus musculus cDNA clone 405277 5') and AA325286 (EST28500 Cerebellum II Homo sapiens cDNA 5' end). The predicted amino acid sequence disclosed herein for hj968\_2 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted hj968\_2 protein demonstrated at least some similarity to the sequence identified as U23528 (translated cosmid B0034.[Caenorhabditis elegans]). Based upon sequence similarity, hj968\_2 proteins and each similar protein or peptide may share at least some activity.

#### Clone "hk10\_3"

A polynucleotide of the present invention has been identified as clone "hk10\_3". hk10\_3 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. hk10\_3 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "hk10\_3 protein").

The nucleotide sequence of hk10\_3 as presently determined is reported in SEQ ID NO:51. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the hk10\_3 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:52. Amino acids 28 to 40 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 41, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone hk10\_3 should be approximately 1300 bp.

The nucleotide sequence disclosed herein for hk10\_3 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. hk10\_3 demonstrated at least some similarity with sequences identified as AA305975 (EST176966 Jurkat T-cells VI Homo sapiens cDNA 5' end  
 5 similar to S. cerevisiae hypothetical protein FAA3-BET1), F18016 (H.sapiens EST sequence 016-T), N36880 (yy37e07.s1 Homo sapiens cDNA clone 273444 3'), and R87757 (yo45a08.s1 Homo sapiens cDNA clone 180854 3'). The predicted amino acid sequence disclosed herein for hk10\_3 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted  
 10 hk10\_3 protein demonstrated at least some similarity to the sequence identified as Z38113 (CAI 0.21 [Saccharomyces cerevisiae]). Based upon sequence similarity, hk10\_3 proteins and each similar protein or peptide may share at least some activity.

#### Clone "hm236\_1"

15 A polynucleotide of the present invention has been identified as clone "hm236\_1". hm236\_1 was isolated from a human adult testes (teratocarcinoma NCCIT) cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded  
 20 protein. hm236\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "hm236\_1 protein").

The nucleotide sequence of hm236\_1 as presently determined is reported in SEQ ID NO:53. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the hm236\_1 protein corresponding to the foregoing  
 25 nucleotide sequence is reported in SEQ ID NO:54.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone hm236\_1 should be approximately 1800 bp.

The nucleotide sequence disclosed herein for hm236\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
 30 FASTA search protocols. hm236\_1 demonstrated at least some similarity with sequences identified as AA026169 (zk01b03.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 469229 5'), AA046211 (zk77e04.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 488862 3'), AA223088 (PMY0368 KG1a Lambda Zap Express cDNA Library Homo sapiens cDNA 5'), AB002368 and AC003010 (Human  
 35 mRNA for KIAA0370 gene, partial cds), AC002399 (Human chromosome 16p11.2 BAC clone CIT987SK-A-481B3; HTGS phase 1, 18 unordered pieces, sequencing in progress), R86676 (ym86f03.r1 Homo sapiens cDNA clone), and T21501 (Human gene signature HUMGS02874). The predicted amino acid sequence disclosed herein for hm236\_1 was

searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted hm236\_1 protein demonstrated at least some similarity to sequences identified as AB002368 (KIAA0370 [Homo sapiens]). Based upon sequence similarity, hm236\_1 proteins and each similar protein or peptide may share at least some activity.

#### Clone "do15\_4"

A polynucleotide of the present invention has been identified as clone "do15\_4". do15\_4 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. do15\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "do15\_4 protein").

The nucleotide sequence of do15\_4 as presently determined is reported in SEQ ID NO:67. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the do15\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:68. Amino acids 394 to 406 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 407, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone do15\_4 should be approximately 1900 bp.

The nucleotide sequence disclosed herein for do15\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. do15\_4 demonstrated at least some similarity with sequences identified as AA113909 (zm80f12.r1 Stratagene neuroepithelium (#937231) Homo sapiens cDNA clone 531983 5'), AA189888 (mu55h06.r1 Soares mouse lymph node NbMLN Mus musculus cDNA clone 643355 5'), and U52052 (Human S6 A-8 mRNA expressed in chromosome 6-suppressed melanoma cells). Based upon sequence similarity, do15\_4 proteins and each similar protein or peptide may share at least some activity.

#### Clone "dx290\_1"

A polynucleotide of the present invention has been identified as clone "dx290\_1". dx290\_1 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. dx290\_1

is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "dx290\_1 protein").

The nucleotide sequence of dx290\_1 as presently determined is reported in SEQ ID NO:69. What applicants presently believe to be the proper reading frame and the  
5 predicted amino acid sequence of the dx290\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:70.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone dx290\_1 should be approximately 2300 bp.

The nucleotide sequence disclosed herein for dx290\_1 was searched against the  
10 GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. dx290\_1 demonstrated at least some similarity with the sequence identified as AA064383 (ml47h02.r1 Stratagene mouse testis (#937308) Mus musculus cDNA clone 515187 5'). Based upon sequence similarity, dx290\_1 proteins and each similar protein or peptide may share at least some activity.

15

#### Clone "ek390\_4"

A polynucleotide of the present invention has been identified as clone "ek390\_4". ek390\_4 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637),  
20 or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ek390\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ek390\_4 protein").

The nucleotide sequence of ek390\_4 as presently determined is reported in SEQ  
25 ID NO:71. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ek390\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:72. Amino acids 25 to 37 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 38, or are a transmembrane domain.

30 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ek390\_4 should be approximately 1000 bp.

The nucleotide sequence disclosed herein for ek390\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ek390\_4 demonstrated at least some similarity with sequences  
35 identified as AA075783 (zm89h02.r1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA clone 545139 5'), AA427538 (zw32g04.r1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 771030 5'), AA427539 (zw32g04.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 771030 3'), AA453353 (zx47a06.r1 Soares testis



NHT Homo sapiens cDNA clone 795346 5'), C20637 (HUMGS0004639, Human Gene Signature, 3'-directed cDNA sequence), R74326 (yl01c07.s1 Homo sapiens cDNA clone 156972 3'), R74420 (yl01c07.r1 Homo sapiens cDNA clone 156972 5'), T22914 (Human gene signature), U41197 (Human [TTTC]10 short tandem repeat polymorphism  
 5 UM65, D17S1340), and X58237 (Human mRNA for anti-lectin antibody epitope (clone p36/8-6)). Based upon sequence similarity, ek390\_4 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the ek390\_4 protein sequence centered around amino acid 160 of SEQ ID NO:72. The nucleotide sequence of ek390\_4  
 10 indicates that it may contain GGGA repeat sequences.

#### Clone "er471\_7"

A polynucleotide of the present invention has been identified as clone "er471\_7". er471\_7 was isolated from a human fetal brain cDNA library using methods which are  
 15 selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. er471\_7 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "er471\_7 protein").

20 The nucleotide sequence of er471\_7 as presently determined is reported in SEQ ID NO:73. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the er471\_7 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:74. Amino acids 74 to 86 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at  
 25 amino acid 87, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone er471\_7 should be approximately 2250 bp.

The nucleotide sequence disclosed herein for er471\_7 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
 30 FASTA search protocols. er471\_7 demonstrated at least some similarity with sequences identified as AA039137 (mi98h06.r1 Soares mouse embryo NbME13.5 14.5 Mus musculus cDNA clone 474683 5'), AA066962 (mm38g05.r1 Stratagene mouse melanoma (#937312) Mus musculus cDNA clone 523832 5'), AA189170 (zq47h05.s1 Stratagene hNT neuron (#937233) Homo sapiens cDNA clone 632889 3'), AA609188  
 35 (af12c10.s1 Soares testis NHT Homo sapiens cDNA clone 1031442 3'), and W07704 (zb02e02.r1 Soares fetal lung NbHL19W Homo sapiens cDNA clone 300890 5' similar to SW:YN66\_YEAST P40164 HYPOTHETICAL 98.1 KD PROTEIN IN SPX19-GCR2 INTERGENIC REGION). The predicted amino acid sequence disclosed herein for

er471\_7 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted er471\_7 protein demonstrated at least some similarity to sequences identified as AF016448 (Cosmid F41E6 [Caenorhabditis elegans]) and L08407 (collagen type XVII [Mus musculus]). Based upon sequence similarity, er471\_7 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts three potential transmembrane domains within the er471\_7 protein sequence, centered around amino acids 40, 80, and 110 of SEQ ID NO:74, respectively.

#### Clone "fs40\_3"

A polynucleotide of the present invention has been identified as clone "fs40\_3". fs40\_3 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fs40\_3 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "fs40\_3 protein").

The nucleotide sequence of fs40\_3 as presently determined is reported in SEQ ID NO:75. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the fs40\_3 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:76.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fs40\_3 should be approximately 1000 bp.

The nucleotide sequence disclosed herein for fs40\_3 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fs40\_3 demonstrated at least some similarity with sequences identified as AA411142 (zt37g01.r1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 724560 5'), AA412527 (zu12a03.s1 Soares testis NHT Homo sapiens cDNA clone 731596 3'), AA565855 (nj32d09.s1 NCI\_CGAP\_AA1 Homo sapiens cDNA clone IMAGE:994193), H17042 (ym39f12.s1 Homo sapiens cDNA clone 50584 3'), and T33280 (EST57284 Homo sapiens cDNA 3' end similar to None). Based upon sequence similarity, fs40\_3 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the fs40\_3 protein sequence at the C-terminus of SEQ ID NO:76.

#### Clone "ga63\_6"

A polynucleotide of the present invention has been identified as clone "ga63\_6". ga63\_6 was isolated from a human adult testes cDNA library using methods which are

selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ga63\_6 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ga63\_6 protein").

The nucleotide sequence of ga63\_6 as presently determined is reported in SEQ ID NO:77. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ga63\_6 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:78. Amino acids 11 to 23 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 24, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ga63\_6 should be approximately 2300 bp.

The nucleotide sequence disclosed herein for ga63\_6 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ga63\_6 demonstrated at least some similarity with sequences identified as AA405433 (zu13h10.r1 Soares testis NHT Homo sapiens cDNA clone 731779 5'similar to TR G474970 G474970 SP32 PRECURSOR), AA406076 (zu67c02.s1 Soares testis NHT Homo sapiens cDNA clone 743042 3' similar to TR:G475021 G475021 SP32 PRECURSOR), AA424694 (zu13h10.s1 Soares testis NHT Homo sapiens cDNA clone 731779 3' similar to TR G475021 G475021 SP32 PRECURSOR; contains element TAR1 repetitive element), D16200 (Pig mRNA for sp32, partial sequence), D16203 (Guinea pig mRNA for sp32, complete cds), and D17573 (Mouse mRNA for proacrosin-binding protein (sp32), complete cds). The predicted amino acid sequence disclosed herein for ga63\_6 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted ga63\_6 protein demonstrated at least some similarity to sequences identified as D16200 (sp32 precursor [*Sus scrofa*]), and D17574 (alternative splicing product for proacrosin-binding protein (sp32) [*Mus musculus*]). The sp32 protein is found in the acrosomal vesicle of sperm, which is involved in egg-sperm fusion in fertilization. This protein is initially synthesized as a 61-kDa precursor protein with a putative signal peptide at the amino terminus. The carboxyl-terminal half of the precursor molecule corresponds to the mature sp32 protein. Thus, sp32 is produced by post-translational modification of the precursor. The binding of sp32 to proacrosin may be involved in packaging the acrosin zymogen into the acrosomal matrix. (Baba *et al.*, 1994, *J. Biol. Chem.* **269** (13): 10133-10140, which is incorporated by reference herein). Based upon sequence similarity, ga63\_6 proteins and each similar protein or peptide may share at least some activity.

Clone "gm335\_4"

A polynucleotide of the present invention has been identified as clone "gm335\_4". gm335\_4 was isolated from a human adult uterus cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. gm335\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "gm335\_4 protein").

The nucleotide sequence of gm335\_4 as presently determined is reported in SEQ ID NO:79. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the gm335\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:80. Amino acids 8 to 20 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 21, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone gm335\_4 should be approximately 800 bp.

The nucleotide sequence disclosed herein for gm335\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. gm335\_4 demonstrated at least some similarity with sequences identified as AA055367 (zf20b05.r1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 377457 5'), AC002389 (Human DNA from chromosome 19 specific cosmid R28461, genomic sequence, complete sequence), W08522 (mb46h10.r1 Soares mouse p3NMF19.5 Mus musculus cDNA clone 332515 5'), and X93916 (S.scrofa mRNA (clone VIB11; expressed sequence tag)). Based upon sequence similarity, gm335\_4 proteins and each similar protein or peptide may share at least some activity.

Clone "hy370\_9"

A polynucleotide of the present invention has been identified as clone "hy370\_9". hy370\_9 was isolated from a human adult trachea cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. hy370\_9 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "hy370\_9 protein").

The nucleotide sequence of hy370\_9 as presently determined is reported in SEQ ID NO:81. What applicants presently believe to be the proper reading frame and the

predicted amino acid sequence of the hy370\_9 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:82. Amino acids 8 to 20 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 21, or are a transmembrane domain.

5           The EcoRI/NotI restriction fragment obtainable from the deposit containing clone hy370\_9 should be approximately 1200 bp.

          The nucleotide sequence disclosed herein for hy370\_9 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. hy370\_9 demonstrated at least some similarity with the  
10       sequence identified as AA763313 (vv89h07.r1 Stratagene mouse skin (#937313) Mus musculus cDNA clone 1229629 5'). Based upon sequence similarity, hy370\_9 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts an additional potential transmembrane domain within the hy370\_9 protein sequence centered around amino acid 140 of SEQ ID NO:82.

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#### Clone "ie47\_4"

          A polynucleotide of the present invention has been identified as clone "ie47\_4". ie47\_4 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was  
20       identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ie47\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ie47\_4 protein").

          The nucleotide sequence of ie47\_4 as presently determined is reported in SEQ  
25       ID NO:83. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ie47\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:84. Amino acids 17 to 29 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 30, or are a transmembrane domain.

30           The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ie47\_4 should be approximately 2300 bp.

          The nucleotide sequence disclosed herein for ie47\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ie47\_4 demonstrated at least some similarity with sequences  
35       identified as AA071953 (mf17h08.r1 Life Tech mouse brain Mus musculus cDNA clone 405375 5' similar to TR G304421 G304421 SILENCER ELEMENT), AA207250 (zq82d05.s1 Stratagene hNT neuron (#937233) Homo sapiens cDNA clone 648105 3' similar to TR G304421 G304421 SILENCER ELEMENT), L14938 (Chicken SCG10

protein mRNA, complete cds), L20260 (Mouse SCG10 gene sequence), R49053 (yg58c05.s1 Homo sapiens cDNA clone 37017 3'), S82024 (SCG10 neuron-specific growth-associated protein/stathmin homolog [human, embryo, mRNA]), T25428 (Human gene signature HUMGS07594, T25428 standard; cDNA to mRNA), W54204 (md04a12.r1 Soares mouse embryo NbME13.5 14.5 Mus musculus cDNA clone 367390 5' similar to SW:SCGB\_XENLA Q09002 SCG10 PROTEIN HOMOLOG A), X71433 (X. laevis SCG10 mRNA), and Z99916 (Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 221G9; HTGS phase 1). The predicted amino acid sequence disclosed herein for ie47\_4 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted ie47\_4 protein demonstrated at least some similarity to sequences identified as L14938 (SCG10 protein [Gallus gallus]) and S82024 (SCG10 neuron-specific growth-associated protein/stathmin homolog [human, embryo, Peptide] [Homo sapiens]). SCG10 protein is considered to be a membrane-bound protein present in neural growth cones and developing neurons (Maucuer *et al.*, 1993, *J. Biol. Chem.* 268: 16420-16429; Stein *et al.*, 1988, *Neuron* 1:463-476; which are incorporated by reference herein). Based upon sequence similarity, ie47\_4 proteins and each similar protein or peptide may share at least some activity.

#### 20      Clone "s195\_10"

A polynucleotide of the present invention has been identified as clone "s195\_10". s195\_10 was isolated from a human adult neural tissue cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. s195\_10 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "s195\_10 protein").

The nucleotide sequence of s195\_10 as presently determined is reported in SEQ ID NO:85. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the s195\_10 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:86. Amino acids 35 to 47 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 48, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone s195\_10 should be approximately 3500 bp.

The nucleotide sequence disclosed herein for s195\_10 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. s195\_10 demonstrated at least some similarity with sequences

identified as AA113800 (zn65b05.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone 563025 3' similar to TR:G600018 G600018 SSM4P), AA114062 (zn65b05.r1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone 563025 5'), AA280316 (zt10f06.s1 Soares NbHTGBC Homo sapiens cDNA clone 712739 3'), AF009301 (Homo sapiens TEB4 protein mRNA, complete cds), N70344 (za60f10.s1 Homo sapiens cDNA clone 296971 3'), R60474 (yh13g07.r1 Homo sapiens cDNA clone 43058 5'), and T26266 (standard; cDNA to mRNA; 148 BP, Human gene signature HUMGS08505). The predicted amino acid sequence disclosed herein for s195\_10 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted s195\_10 protein demonstrated at least some similarity to sequences identified as AF009301 (TEB4 protein [Homo sapiens]), X76715 (SSM4 gene product [Saccharomyces cerevisiae]), Z46861 (Ssm4p [Saccharomyces cerevisiae]), and Z47047 (Ssm4p [Saccharomyces cerevisiae]). Based upon sequence similarity, s195\_10 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts eleven additional potential transmembrane domains within the s195\_10 protein sequence, centered around amino acids 130, 170, 210, 260, 320, 470, 520, 560, 600, 650, and 690 of SEQ ID NO:86, respectively. The nucleotide sequence of s195\_10 indicates that it may contain a simple GAA repeat region.

#### Clone "bf228\_14"

A polynucleotide of the present invention has been identified as clone "bf228\_14". bf228\_14 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bf228\_14 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bf228\_14 protein").

The nucleotide sequence of bf228\_14 as presently determined is reported in SEQ ID NO:97. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bf228\_14 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:98. Amino acids 18 to 30 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 31, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bf228\_14 should be approximately 1400 bp.

The nucleotide sequence disclosed herein for bf228\_14 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and

FASTA search protocols. bf228\_14 demonstrated at least some homology with sequences identified as AA069549 (zm52e03.s1 Stratagene fibroblast (#937212) Homo sapiens cDNA clone 529276 3'), H83278 (yq49h09.r1 Homo sapiens cDNA clone 199169 5'), and N94898 (zb31b01.s1 Homo sapiens cDNA clone 305161 3' similar to contains Alu repetitive element; contains element MSR1 repetitive element). Based upon homology, bf228\_14 proteins and each homologous protein or peptide may share at least some activity. The nucleotide sequence of bf228\_14 indicates that it may contain an Alu repetitive element.

#### 10 Clone "bg249\_1"

A polynucleotide of the present invention has been identified as clone "bg249\_1". bg249\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bg249\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bg249\_1 protein").

The nucleotide sequence of bg249\_1 as presently determined is reported in SEQ ID NO:99. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bg249\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:100.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bg249\_1 should be approximately 2700 bp.

The nucleotide sequence disclosed herein for bg249\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bg249\_1 demonstrated at least some homology with sequences identified as AA151021 (zl47c04.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 505062 5'), AA278781 (zs79a01.r1 Soares NbHTGBC Homo sapiens cDNA clone 703656 5'), R75099 (MDB1032R Mouse brain, Stratagene Mus musculus cDNA 5'end), and T06990 (EST04879 Homo sapiens cDNA clone HFBEB91). Based upon homology, bg249\_1 proteins and each homologous protein or peptide may share at least some activity.

#### 35 Clone "bv286\_1"

A polynucleotide of the present invention has been identified as clone "bv286\_1". bv286\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the



basis of computer analysis of the amino acid sequence of the encoded protein. bv286\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bv286\_1 protein").

The nucleotide sequence of bv286\_1 as presently determined is reported in SEQ ID NO:101. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bv286\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:102. Amino acids 14 to 26 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 27, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bv286\_1 should be approximately 550 bp.

The nucleotide sequence disclosed herein for bv286\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bv286\_1 demonstrated at least some homology with sequences identified as AA132163 (zl38c07.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 504204 5' similar to WP:F55A11.1 CE05943 EF HAND DOMAINS), AA552888 (nk57h08.s1 NCI\_CGAP\_Pr7 Homo sapiens cDNA clone IMAGE:1017663 similar to WP:F55A11.1 CE05943 EF HAND DOMAINS), and H12316 (yj11d07.s1 Homo sapiens cDNA clone 148429 3' similar to SP:JS0027 JS0027 PROBABLE CALCIUM-BINDING PROTEIN). The predicted amino acid sequence disclosed herein for bv286\_1 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bv286\_1 protein demonstrated at least some identity with sequences identified as L22647 (prostaglandin receptor ep1 subtype [Homo sapiens]). Based upon homology, bv286\_1 proteins and each homologous protein or peptide may share at least some activity. The nucleotide sequence of bv286\_1 indicates that it may contain an Alu repetitive element.

#### Clone "co36\_1"

A polynucleotide of the present invention has been identified as clone "co36\_1". co36\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. co36\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "co36\_1 protein").

The nucleotide sequence of co36\_1 as presently determined is reported in SEQ ID NO:103. What applicants presently believe to be the proper reading frame and the

predicted amino acid sequence of the co36\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:104.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone co36\_1 should be approximately 3300 bp.

5 The nucleotide sequence disclosed herein for co36\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. No hits were found in the database.

# 10 Clone "cp116\_1"

A polynucleotide of the present invention has been identified as clone "cp116\_1". cp116\_1 was isolated from a human adult salivary gland cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cp116\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cp116\_1 protein").

The nucleotide sequence of cp116\_1 as presently determined is reported in SEQ ID NO:105. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cp116\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:106. Amino acids 3 to 15 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 16, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cp116\_1 should be approximately 1600 bp.

The nucleotide sequence disclosed herein for cp116\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. No hits were found in the database.

# 30 Clone "cw1195\_2"

A polynucleotide of the present invention has been identified as clone "cw1195\_2". cw1195\_2 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cw1195\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cw1195\_2 protein").

The nucleotide sequence of the 5' portion of cw1195\_2 as presently determined is reported in SEQ ID NO:107. An additional internal nucleotide sequence from cw1195\_2 as presently determined is reported in SEQ ID NO:108. What applicants believe is the proper reading frame and the predicted amino acid sequence encoded by such internal sequence is reported in SEQ ID NO:109. Additional nucleotide sequence from the 3' portion of cw1195\_2, including the polyA tail, is reported in SEQ ID NO:110.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cw1195\_2 should be approximately 3300 bp.

The nucleotide sequence disclosed herein for cw1195\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cw1195\_2 demonstrated at least some homology with sequences identified as AA205460 (zq66f07.s1 Stratagene neuroepithelium (#937231) Homo sapiens cDNA clone 646597 3') and AA362052 (EST71451 MCF7 cell line Homo sapiens cDNA 5' end similar to EST containing Alu repeat). Based upon homology, cw1195\_2 proteins and each homologous protein or peptide may share at least some activity. The nucleotide sequence of cw1195\_2 indicates that it may contain an Alu repetitive element.

#### Clone "fh13\_10"

A polynucleotide of the present invention has been identified as clone "fh13\_10". fh13\_10 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. fh13\_10 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "fh13\_10 protein").

The nucleotide sequence of fh13\_10 as presently determined is reported in SEQ ID NO:111. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the fh13\_10 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:112.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone fh13\_10 should be approximately 1400 bp.

The nucleotide sequence disclosed herein for fh13\_10 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. fh13\_10 demonstrated at least some homology with sequences identified as J00089 (Human Alu family interspersed repeat; clone BLUR6) and X00481 (Human non-alu family interspersed repeat). Based upon homology, fh13\_10 proteins and each homologous protein or peptide may share at least some

activity. The nucleotide sequence of fh13\_10 indicates that it may contain a repetitive element.

5           Clone "gc57\_4"

A polynucleotide of the present invention has been identified as clone "gc57\_4". gc57\_4 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. gc57\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "gc57\_4 protein").

The nucleotide sequence of gc57\_4 as presently determined is reported in SEQ ID NO:113. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the gc57\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:114. Amino acids 10 to 22 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 23, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone gc57\_4 should be approximately 1900 bp.

The nucleotide sequence disclosed herein for gc57\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. gc57\_4 demonstrated at least some homology with sequences identified as AA095328 (I3005.seq.F Fetal heart, Lambda ZAP Express Homo sapiens cDNA 5'), AA126440 (zk94e02.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 490490 3'), and Z82195 (Human DNA sequence from clone J274L71). Based upon homology, gc57\_4 proteins and each homologous protein or peptide may share at least some activity. The nucleotide sequence of gc57\_4 indicates that it may contain an Alu repetitive element.

30

Clone "h1165\_3"

A polynucleotide of the present invention has been identified as clone "h1165\_3". h1165\_3 was isolated from a human adult blood (peripheral blood mononuclear cells treated with phytohemagglutinin and phorbol meristate acetate and mixed lymphocyte reaction) cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. h1165\_3 is a full-length clone, including

the entire coding sequence of a secreted protein (also referred to herein as "h1165\_3 protein").

The nucleotide sequence of h1165\_3 as presently determined is reported in SEQ ID NO:115. What applicants presently believe to be the proper reading frame and the  
5 predicted amino acid sequence of the h1165\_3 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:116.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone h1165\_3 should be approximately 1250 bp.

The nucleotide sequence disclosed herein for h1165\_3 was searched against the  
10 GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. h1165\_3 demonstrated at least some homology with sequences identified as AA173098 (zp31d03.r1 Stratagene neuroepithelium (#937231) Homo sapiens cDNA clone 611045 5' similar to contains element TAR1 repetitive element), AA305139 (EST176159 Colon carcinoma (Caco-2) cell line II Homo sapiens  
15 cDNA 5' end), AA426375 (zv54h02.s1 Soares testis NHT Homo sapiens cDNA clone 757491 3'), and N72370 (yv38c11.r1 Homo sapiens cDNA clone 245012 5'). The predicted amino acid sequence disclosed herein for h1165\_3 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted h1165\_3 protein demonstrated at least some identity with  
20 sequences identified as U58758 (coded for by C. elegans cDNA yk83a5.3; coded for by C.elegans cDNA yk83a5.5 [Caenorhabditis elegans]). Based upon homology, h1165\_3 proteins and each homologous protein or peptide may share at least some activity.

#### Clone "hb752\_1"

25 A polynucleotide of the present invention has been identified as clone "hb752\_1". hb752\_1 was isolated from a human fetal kidney cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. hb752\_1  
30 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "hb752\_1 protein").

The nucleotide sequence of hb752\_1 as presently determined is reported in SEQ ID NO:117. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the hb752\_1 protein corresponding to the foregoing  
35 nucleotide sequence is reported in SEQ ID NO:118. Amino acids 16 to 28 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 29, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone hb752\_1 should be approximately 1800 bp.

The nucleotide sequence disclosed herein for hb752\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. hb752\_1 demonstrated at least some homology with sequences identified as AA100979 (zm26f09.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone 526793 3'), AA490528 (aa51g11.s1 NCI\_CGAP\_GCB1 Homo sapiens cDNA clone 824516 3'), H65670 (yr72g12.r1 Homo sapiens cDNA clone 210886 5'), H86790 (ys72c03.s1 Homo sapiens cDNA clone 220324 3'), N58917 (yy61f11.s1 Homo sapiens cDNA clone 278061 3'), and Z43307 (H. sapiens partial cDNA sequence; clone c-18g09). Based upon homology, hb752\_1 proteins and each homologous protein or peptide may share at least some activity.

#### Clone "bi127\_5"

A polynucleotide of the present invention has been identified as clone "bi127\_5". bi127\_5 was isolated from a human fetal kidney cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bi127\_5 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bi127\_5 protein").

The nucleotide sequence of bi127\_5 as presently determined is reported in SEQ ID NO:129. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bi127\_5 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:130.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bi127\_5 should be approximately 2500 bp.

The nucleotide sequence disclosed herein for bi127\_5 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bi127\_5 demonstrated at least some similarity with sequences identified as AA055840 (zf20c06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 377482 3'), AA334304 (EST38496 Embryo, 9 week Homo sapiens cDNA 5' end similar to similar to H. sapiens hypothetical protein, chromosome 3p21.1 gene sequence (GB:L13435)), AA399397 (zt59f11.r1 Soares testis NHT Homo sapiens cDNA clone 726669 5'), AA576692 (nm73a07.s1 NCI\_CGAP\_Co9 Homo sapiens cDNA clone IMAGE:1073844), H01918 (yj29a07.s1 Homo sapiens cDNA clone 150132 3'), H94897 (yu57h08.s1 Homo sapiens cDNA clone 230271 3'), L13435 (Human chromosome 3p21.1 gene sequence), R85965 (yt66g02.s1 Soares retina N2b4HR Homo sapiens

cDNA clone 275499 3'), and X95828 (H.sapiens DNA NotI jumping clone J32A032D). Based upon sequence similarity, bi127\_5 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the bi127\_5 protein sequence centered around amino  
 5 acid 15 of SEQ ID NO:130.

#### Clone "bl194\_2"

A polynucleotide of the present invention has been identified as clone "bl194\_2". bl194\_2 was isolated from a human adult testes cDNA library using methods which are  
 10 selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bl194\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bl194\_2 protein").

15 The nucleotide sequence of bl194\_2 as presently determined is reported in SEQ ID NO:130. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bl194\_2 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:132. Amino acids 88 to 100 are a predicted leader/signal sequence, with the predicted mature amino acid sequence  
 20 beginning at amino acid 101, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bl194\_2 should be approximately 1600 bp.

The nucleotide sequence disclosed herein for bl194\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
 25 FASTA search protocols. bl194\_2 demonstrated at least some similarity with sequences identified as AA136931 (zn97f05.s1 Stratagene fetal retina 937202 Homo sapiens cDNA clone 566145 3'), AA148976 (zn99e10.r1 Stratagene colon (#937204) Homo sapiens cDNA clone 566346 5'), AA148977 (zn99e10.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 566346 3'), AA196293 (zp92g07.s1 Stratagene HeLa cell s3  
 30 937216 Homo sapiens cDNA clone 627708 3'), AA487754 (ab13e12.r1 Stratagene lung (#937210) Homo sapiens cDNA clone 840718 5'), H01254 (yj27b02.r1 Homo sapiens cDNA clone 149931 5'), H86324 (yt05f07.r1 Homo sapiens cDNA clone 223429 5'), N23958 (yx71c02.s1 Homo sapiens cDNA clone 267170 3'), N31859 (yx71c02.r1 Homo sapiens cDNA clone 267170 5'), R01674 (ye76b07.s1 Homo sapiens cDNA clone  
 35 123637 3'), and T78480 (yd68g08.s1 Homo sapiens cDNA clone 113438 3'). The predicted amino acid sequence disclosed herein for bl194\_2 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bl194\_2 protein demonstrated at least some similarity to the

sequence identified as S75895 (NADH dehydrogenase subunit 2, ND2 [human, brain, Peptide Mitochondrial Partial Mutant, 79 aa] [Homo sapiens]). Based upon sequence similarity, bl194\_2 proteins and each similar protein or peptide may share at least some activity.

5

Clone "cc130\_1"

A polynucleotide of the present invention has been identified as clone "cc130\_1". cc130\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was  
10 identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cc130\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cc130\_1 protein").

The nucleotide sequence of cc130\_1 as presently determined is reported in SEQ  
15 ID NO:133. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cc130\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:134. Amino acids 7 to 19 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 20, or are a transmembrane domain.

20 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cc130\_1 should be approximately 2500 bp.

The nucleotide sequence disclosed herein for cc130\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cc130\_1 demonstrated at least some similarity with sequences  
25 identified as T21181 (Human gene signature HUMGS02491), T70127 (yc17d06.r1 Homo sapiens cDNA clone 80939 5' similar to SP:BUTY\_BOVIN P18892 BUTYROPHILIN PRECURSOR), T92875 (ye27h03.r1 Homo sapiens cDNA clone 118997 5'), U39576 (Human butyrophilin precursor mRNA, complete cds), U90546 (Human butyrophilin (BTF4) mRNA, complete cds), and W69453 (zd45e02.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 343610 3'). The predicted amino acid sequence disclosed herein for cc130\_1 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted cc130\_1 protein demonstrated at least some similarity to sequences identified  
30 as M35551 (BOVBUTBT1\_1 Bovine butyrophilin mRNA, complete cds. [Bos taurus]), R71361 (Human truncated MOG), U39576 (butyrophilin precursor [Homo sapiens]), and U90546 (butyrophilin [Homo sapiens]). Butyrophilin may function in the secretion of milk-fat droplets and may act as a specific membrane-associated receptor for the association of cytoplasmic droplets with the apical plasma membrane. The subcellular



location of butyrophilin is that of a Type I membrane protein. Butyrophilin also exhibits tissue specificity, being expressed in mammary tissue and secreted in association with the milk-fat-globule membrane during lactation. Butyrophilin is also homologous to MOG (myelinoligo dendrocyte protein) which is used to treat auto-immune diseases.

5 Both butyrophilin and MOG are homologous in the same amino acids to an immunoglobulin variable region; this may indicate the existence of a protein-protein binding (receptor) site. Based upon sequence similarity, cc130\_1 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts an additional potential transmembrane domain within the cc130\_1

10 protein sequence centered around amino acid 255 of SEQ ID NO:134.

#### Clone "ch582\_1"

A polynucleotide of the present invention has been identified as clone "ch582\_1". ch582\_1 was isolated from a human fetal kidney cDNA library using

15 methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ch582\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ch582\_1 protein").

20 The nucleotide sequence of ch582\_1 as presently determined is reported in SEQ ID NO:135. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ch582\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:136. Amino acids 23 to 35 are a predicted leader/signal sequence, with the predicted mature amino acid sequence

25 beginning at amino acid 36, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ch582\_1 should be approximately 2300 bp.

The nucleotide sequence of ch582\_1 indicates that it may contain one or more repetitive elements.

30

#### Clone "cq294\_14"

A polynucleotide of the present invention has been identified as clone "cq294\_14". cq294\_14 was isolated from a human adult heart cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No.

35 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cq294\_14 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cq294\_14 protein").

The nucleotide sequence of cq294\_14 as presently determined is reported in SEQ ID NO:137. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cq294\_14 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:138.

5 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cq294\_14 should be approximately 1850 bp.

The nucleotide sequence disclosed herein for cq294\_14 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cq294\_14 demonstrated at least some similarity with  
 10 sequences identified as AA133962 (zl34c12.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 503830 3'), AA447968 (zv83h10.s1 Soares total fetus Nb2HF8 9w Homo sapiens cDNA clone 760291 3'), N47086 (yy85c08.s1 Homo sapiens cDNA clone 280334 3'), R33663 (yh82g06.s1 Homo sapiens cDNA clone 136282 3'), R45544 (yg43g12.s1 Homo sapiens cDNA clone 35358 3'), R77637 (yi76h09.s1 Homo sapiens  
 15 cDNA clone 145217 3'), and W37736 (zc10h10.r1 Soares parathyroid tumor NbHPA Homo sapiens cDNA clone). Based upon sequence similarity, cq294\_14 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts three potential transmembrane domains within the cq294\_14 protein sequence, centered around amino acids 15, 25, and 50 of SEQ ID  
 20 NO:138, respectively. The nucleotide sequence of cq294\_14 indicates that it may contain one or more repetitive elements.

#### Clone "dd454\_1"

A polynucleotide of the present invention has been identified as clone  
 25 "dd454\_1". dd454\_1 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. dd454\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also  
 30 referred to herein as "dd454\_1 protein").

The nucleotide sequence of dd454\_1 as presently determined is reported in SEQ ID NO:139. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the dd454\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:140.

35 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone dd454\_1 should be approximately 2300 bp.

The nucleotide sequence disclosed herein for dd454\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and

FASTA search protocols. dd454\_1 demonstrated at least some similarity with sequences identified as AA393499 (zt73g06.r1 Soares testis NHT Homo sapiens cDNA clone 728026 5') and AA430063 (zw67a12.s1 Soares testis NHT Homo sapiens cDNA clone 781246 3'). Based upon sequence similarity, dd454\_1 proteins and each similar  
5 protein or peptide may share at least some activity.

Clone "du157\_12"

A polynucleotide of the present invention has been identified as clone "du157\_12". du157\_12 was isolated from a human fetal brain cDNA library using  
10 methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. du157\_12 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "du157\_12 protein").

15 The nucleotide sequence of du157\_12 as presently determined is reported in SEQ ID NO:141. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the du157\_12 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:142.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
20 clone du157\_12 should be approximately 4050 bp.

The nucleotide sequence disclosed herein for du157\_12 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. du157\_12 demonstrated at least some similarity with sequences identified as AA164862 (zq41g04.r1 Stratagene hNT neuron (#937233) Homo sapiens cDNA clone 632310 5'), AA284379 (zs59a07.r1 NCI\_CGAP\_GCB1 Homo sapiens cDNA clone IMAGE:701748 5'), and T22493 (Human gene signature HUMGS04104). Based upon sequence similarity, du157\_12 proteins and each similar  
25 protein or peptide may share at least some activity.

30 Clone "du372\_1"

A polynucleotide of the present invention has been identified as clone "du372\_1". du372\_1 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the  
35 basis of computer analysis of the amino acid sequence of the encoded protein. du372\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "du372\_1 protein").

The nucleotide sequence of du372\_1 as presently determined is reported in SEQ ID NO:143. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the du372\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:145. Amino acids 69 to 81 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 82, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone du372\_1 should be approximately 1500 bp.

The nucleotide sequence disclosed herein for du372\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. du372\_1 demonstrated at least some similarity with sequences identified as AA099051 (zn45c07.r1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone 550380 5'), AA424986 (zw06g01.r1 Soares NhHMPu S1 Homo sapiens cDNA clone 768528 5'), AA480114 (zv41h05.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 756249 3'), H73153 (yu26e11.r1 Homo sapiens cDNA clone 234956 5'), and H73629 (yu26f11.r1 Homo sapiens cDNA clone 234957 5'). Based upon sequence similarity, du372\_1 proteins and each similar protein or peptide may share at least some activity.

#### Clone "ej90\_5"

A polynucleotide of the present invention has been identified as clone "ej90\_5". ej90\_5 was isolated from a human adult placenta cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ej90\_5 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ej90\_5 protein").

The nucleotide sequence of ej90\_5 as presently determined is reported in SEQ ID NO:145. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ej90\_5 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:146. Amino acids 8 to 20 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 21, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ej90\_5 should be approximately 850 bp.

The nucleotide sequence disclosed herein for ej90\_5 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ej90\_5 demonstrated at least some similarity with sequences

identified as AA099387 (zk85e10.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 489642 5'), AA099388 (zk85e10.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 489642 3'), AA256657 (zr85c06.r1 Soares NhHMPu S1 Homo sapiens cDNA clone 682474 5'), and X85111 (X.laevis mRNA for XEL-1). Based upon  
 5 sequence similarity, ej90\_5 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts an additional potential trans-membrane domain within the ej90\_5 protein sequence centered around amino acid 164 of SEQ ID NO:146.

10 Clone "ic2\_6"

A polynucleotide of the present invention has been identified as clone "ic2\_6". ic2\_6 was isolated from a human adult retina (retinoblastoma WERI-Rb1) cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on  
 15 the basis of computer analysis of the amino acid sequence of the encoded protein. ic2\_6 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ic2\_6 protein").

The nucleotide sequence of ic2\_6 as presently determined is reported in SEQ ID NO:147. What applicants presently believe to be the proper reading frame and the  
 20 predicted amino acid sequence of the ic2\_6 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:148. Amino acids 5 to 17 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 18, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
 25 clone ic2\_6 should be approximately 2000 bp.

The nucleotide sequence disclosed herein for ic2\_6 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ic2\_6 demonstrated at least some similarity with sequences  
 30 identified as AA104139 (mp03a12.r1 Stratagene mouse heart (#937316) Mus musculus cDNA clone 568126 5'), N39195 (yv26e08.s1 Homo sapiens cDNA clone 243878 3'), and Z59762 (H.sapiens CpG DNA, clone 171h5, reverse read cpg171h5.rt1a). Based upon sequence similarity, ic2\_6 proteins and each similar protein or peptide may share at least some activity.

35 Clone "bn97\_1"

A polynucleotide of the present invention has been identified as clone "bn97\_1". bn97\_1 was isolated from a human adult placenta cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the

amino acid sequence of the encoded protein. bn97\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bn97\_1 protein").

5 The nucleotide sequence of bn97\_1 as presently determined is reported in SEQ ID NO:159. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bn97\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:160. Amino acids 55 to 67 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 68, or are a transmembrane domain.

10 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bn97\_1 should be approximately 1700 bp.

The nucleotide sequence disclosed herein for bn97\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bn97\_1 demonstrated at least some identity with sequences  
 15 identified as AA046689 (zk72h06.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 488411 3'), D30934 (Human fetal-lung cDNA 5'-end sequence), R78820 (yi90b03.r1 Homo sapiens cDNA clone 146477 5'), and R91687 (yq10h09.s1 Homo sapiens cDNA clone 196577 3'). The predicted amino acid sequence disclosed herein for bn97\_1 was searched against the GenPept and GeneSeq amino acid sequence  
 20 databases using the BLASTX search protocol. The predicted bn97\_1 protein demonstrated at least some identity with sequences identified as A10431 (Hepatitis-B virus surface antigen P31). The bn97\_1 protein also shows some identity (30% identity, 50% conserved amino acids) to both bovine and human lectin-like receptor for oxidized LDL (low-density lipoprotein). While this homology is weak, it gets stronger (44%  
 25 identity and 62% conserved amino acids) in the lectin-like domain. Further, the 3' untranslated region of the bovine receptor has seven mRNA unstabilising sequences (ATTTA) and bn97\_1 has four in its 3' untranslated region. This lectin-like receptor for oxidized LDL (designated LOX-1, Sawamura *et al.*, 1997, *Nature* 386: 73-77) is an integral membrane protein which binds oxidized low-density lipoproteins, internalizes  
 30 them into the endothelial cells and destroys them, thus playing a crucial role in the pathogenesis of atherosclerosis. Based upon identity, bn97\_1 proteins and each identical protein or peptide may share at least some activity.

#### Clone "bn268\_11"

35 A polynucleotide of the present invention has been identified as clone "bn268\_11". bn268\_11 was isolated from a human adult placenta cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bn268\_11 is a full-length

clone, including the entire coding sequence of a secreted protein (also referred to herein as "bn268\_11 protein").

The nucleotide sequence of bn268\_11 as presently determined is reported in SEQ ID NO:161. What applicants presently believe to be the proper reading frame and  
5 the predicted amino acid sequence of the bn268\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:162.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bn268\_11 should be approximately 1050 bp.

The nucleotide sequence disclosed herein for bn268\_11 was searched against the  
10 GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bn268\_11 demonstrated at least some identity with sequences identified as D62832 (Human aorta cDNA 5'-end GEN-330C09) and U20159 (Mus musculus 76 kDa tyrosine phosphoprotein SLP-76 mRNA, complete cds). The predicted amino acid sequence disclosed herein for bn268\_11 was searched against the  
15 GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bn268\_11 protein demonstrated at least some identity with sequences identified as D83171 (GDP-GTP exchange protein for Rho1p [Saccharomyces cerevisiae]). Based upon identity, bn268\_11 proteins and each identical protein or peptide may share at least some activity. The TopPredII computer  
20 program predicts a potential transmembrane domain within the bn268\_11 protein sequence centered around amino acid 33 of SEQ ID NO:162; this region may also function as a signal sequence.

#### Clone "cb96\_10"

25 A polynucleotide of the present invention has been identified as clone "cb96\_10". cb96\_10 was isolated from a human fetal brain cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cb96\_10 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as  
30 "cb96\_10 protein").

The nucleotide sequence of cb96\_10 as presently determined is reported in SEQ ID NO:163. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cb96\_10 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:164. Amino acids 74 to 86 are a  
35 predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 87, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cb96\_10 should be approximately 2100 bp.

The nucleotide sequence disclosed herein for cb96\_10 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cb96\_10 demonstrated at least some identity with sequences identified as AA459012, AA459236, AA256744 (zs31h11.r1 Soares NbHTGBC Homo sapiens cDNA clone 686853 5'), N54489 (yv40f07.s1 Soares fetal liver spleen 1NFLS Homo sapiens cDNA clone 245221 3'), and N57339 (yw81h07.r1 Homo sapiens cDNA clone 258685 5'). The predicted amino acid sequence disclosed herein for cb96\_10 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted cb96\_10 protein demonstrated at least some identity with sequences identified as X80036 (ascorbate peroxidase [Arabidopsis thaliana]). Based upon identity, cb96\_10 proteins and each identical protein or peptide may share at least some activity. The TopPredII computer program predicts seven potential transmembrane domains within the cb96\_10 protein sequence, centered around amino acid residues 25, 80, 125, 225, 300, 350, and 440 of SEQ ID NO:162. Therefore, cb96\_10 is likely to be an integral membrane protein with multiple helices in the membrane; it also contains the sequence motif of the actinin-type actin-binding domains that are believed to anchor actin to the cell membrane.

#### Clone "cb213\_11"

A polynucleotide of the present invention has been identified as clone "cb213\_11". cb213\_11 was isolated from a human fetal brain cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cb213\_11 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cb213\_11 protein").

The nucleotide sequence of cb213\_11 as presently determined is reported in SEQ ID NO:165. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cb213\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:166. Amino acids 29 to 41 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 42, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cb213\_11 should be approximately 2400 bp.

The nucleotide sequence disclosed herein for cb213\_11 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cb213\_11 demonstrated at least some identity with sequences identified as AA332165 (EST36344 Embryo, 8 week I Homo sapiens cDNA 5' end) and R34507 (g58a03.r1 Homo sapiens cDNA clone 36801 5'). The predicted amino acid



sequence disclosed herein for cb213\_11 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted cb213\_11 protein demonstrated at least some identity with sequences identified as U39847 (AO13 ankyrin [Caenorhabditis elegans]). Based upon identity, cb213\_11 proteins and each identical protein or peptide may share at least some activity.

#### Clone "cj457\_4"

A polynucleotide of the present invention has been identified as clone "cj457\_4". cj457\_4 was isolated from a human fetal brain cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cj457\_4 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cj457\_4 protein").

The nucleotide sequence of cj457\_4 as presently determined is reported in SEQ ID NO:167. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cj457\_4 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:168. Amino acids 11 to 23 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 24, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cj457\_4 should be approximately 3350 bp.

The nucleotide sequence disclosed herein for cj457\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cj457\_4 demonstrated at least some identity with sequences identified as T92881 (ye22a10.s1 Homo sapiens cDNA clone 118458 3') and T92488 (ye21g09.r1 Homo sapiens cDNA clone 118432 5'). Based upon identity, cj457\_4 proteins and each identical protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the cj457\_4 protein sequence, centered around amino acid 17 of SEQ ID NO:168; this region may also function as a signal sequence.

#### Clone "cz653\_11"

A polynucleotide of the present invention has been identified as clone "cz653\_11". cz653\_11 was isolated from a human adult testes cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cz653\_11 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cz653\_11 protein").

The nucleotide sequence of cz653\_11 as presently determined is reported in SEQ ID NO:169. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cz653\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:170.

5           The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cz653\_11 should be approximately 1300 bp.

          The nucleotide sequence disclosed herein for cz653\_11 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cz653\_11 demonstrated at least some identity with sequences  
 10 identified as AA024740 (ze76c09.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 364912 3'), AA203204 (zx57b04.r1 Soares fetal liver spleen 1NFLS S1 Homo sapiens cDNA clone 446575 5' similar to contains element MSR1 repetitive element), W72894 (zd59e06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 344962 3'), and W76099 (zd59e06.r1 Soares fetal heart NbHH19W Homo sapiens cDNA clone  
 15 344962 5'). The predicted cz653\_11 demonstrated similarity to various WD-40 repeat containing proteins such as beta transducin-like protein (L28125) and coatomer, beta-prime subunit (AJ006523). The homology appears to be due to the presence of the Beta-transducin family Trp-Asp repeats signature (WD-40) beginning at residue 262 of SEQ ID NO:117. The WD-40 repeat has been thought to be a protein-protein interaction  
 20 domain. Based upon identity, cz653\_11 proteins and each identical protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the cz653\_11 protein sequence centered around amino acid 200 of SEQ ID NO:170.

25           Clone "dx138\_4"

          A polynucleotide of the present invention has been identified as clone "dx138\_4". dx138\_4 was isolated from a human adult testes cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. dx138\_4 is a full-length  
 30 clone, including the entire coding sequence of a secreted protein (also referred to herein as "dx138\_4 protein").

          The nucleotide sequence of dx138\_4 as presently determined is reported in SEQ ID NO:171. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the dx138\_4 protein corresponding to the foregoing  
 35 nucleotide sequence is reported in SEQ ID NO:172. Amino acids 268 to 280 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 281, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone dx138\_4 should be approximately 2300 bp.

The nucleotide sequence disclosed herein for dx138\_4 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. dx138\_4 demonstrated at least some identity with sequences identified as AA108970 (ml63a06.r1 Stratagene mouse testis (#937308) Mus musculus cDNA clone 516658 5'), AA280976 (zs97f01.r1 Soares NbHTGBC Homo sapiens cDNA clone 711577 5' similar to contains Alu repetitive element), H99316 (yx23a03.s1 Homo sapiens cDNA clone 262540 3'), T36050 (EST96120 Homo sapiens cDNA 5'), X85637 (H.sapiens mRNA for expressed sequence tag, clone CAM tEST417 (A)), and Z22280 (H.sapiens DNA sequence). Based upon identity, dx138\_4 proteins and each identical protein or peptide may share at least some activity.

#### Clone "ij167\_5"

A polynucleotide of the present invention has been identified as clone "ij167\_5". ij167\_5 was isolated from a human adult blood (peripheral blood mononuclear cells treated *in vivo* with G-CSF) cDNA library was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ij167\_5 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ij167\_5 protein").

The nucleotide sequence of ij167\_5 as presently determined is reported in SEQ ID NO:173. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ij167\_5 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:174. Amino acids 12 to 24 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 25, or are a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ij167\_5 should be approximately 1050 bp.

The nucleotide sequence disclosed herein for ij167\_5 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ij167\_5 demonstrated at least some identity with sequences identified as N71115 (za87h10.s1 Homo sapiens cDNA clone 299587 3'), T85491 (yd78b01.r1 Homo sapiens cDNA clone 114313 5'), W04374 (za43f06.r1 Soares fetal liver spleen 1NFLS Homo sapiens cDNA clone 295331 5'), W05476 (za87h10.r1 Soares fetal lung NbHL19W Homo sapiens cDNA clone 299587 5'), and W40146 (zb74d09.r1 Soares fetal lung NbHL19W Homo sapiens cDNA clone 309329 5'). The predicted amino acid sequence disclosed herein for ij167\_5 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The

predicted ij167\_5 protein demonstrated at least some identity with sequences identified as M96653 (adenylyl cyclase, type 6 [Mus musculus]). Based upon identity, ij167\_5 proteins and each identical protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the  
 5 ij167\_5 protein sequence, centered around amino acid 40 of SEQ ID NO:174.

#### Clone "bd107\_16"

A polynucleotide of the present invention has been identified as clone "bd107\_16". bd107\_16 was isolated from a human fetal kidney cDNA library using  
 10 methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bd107\_16 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bd107\_16 protein").

15 The nucleotide sequence of bd107\_16 as presently determined is reported in SEQ ID NO:183. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bd107\_16 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:184.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
 20 clone bd107\_16 should be approximately 1500 bp.

The nucleotide sequence disclosed herein for bd107\_16 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bd107\_16 demonstrated at least some similarity with sequences identified as AA261841 (zs17h09.r1 NCI\_CGAP\_GCB1 Homo sapiens cDNA  
 25 clone IMAGE:685505 5'), AA424094 (zv80d05.r1 Soares total fetus Nb2HF8 9w Homo sapiens cDNA clone 759945 5'), AA449717 (zx09b06.s1 Soares total fetus Nb2HF8 9w Homo sapiens cDNA clone 785939 3' similar to TR:E246888 E246888 CHROMOSOME XVI READING FRAME ORF YPL146C), AA875866 (ob34d08.s1 NCI\_CGAP\_Kid5 Homo sapiens cDNA clone IMAGE:1325583 3' similar to TR:Q12080 Q12080 P2610),  
 30 H80410 (yu97b09.r1 Homo sapiens cDNA clone 241145 5'), N39747 (yx92h07.r1 Homo sapiens cDNA clone 269245 5'), R97655 (yq59d12.r1 Homo sapiens cDNA clone 200087 5'), T19822 (Human gene signature HUMGS00904; standard; cDNA to mRNA), and W68551 (zd36h03.r1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 342773 5'). The predicted amino acid sequence disclosed herein for bd107\_16 was searched  
 35 against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bd107\_16 protein demonstrated at least some similarity to the sequence identified as U43703 (Lpi2p [Saccharomyces cerevisiae]). Based upon

sequence similarity, bd107\_16 proteins and each similar protein or peptide may share at least some activity.

Clone "bm41\_7"

5 A polynucleotide of the present invention has been identified as clone "bm41\_7".  
bm41\_7 was isolated from a human adult muscle cDNA library using methods which  
are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was  
identified as encoding a secreted or transmembrane protein on the basis of computer  
analysis of the amino acid sequence of the encoded protein. bm41\_7 is a full-length  
10 clone, including the entire coding sequence of a secreted protein (also referred to herein  
as "bm41\_7 protein").

The nucleotide sequence of bm41\_7 as presently determined is reported in SEQ  
ID NO:185. What applicants presently believe to be the proper reading frame and the  
predicted amino acid sequence of the bm41\_7 protein corresponding to the foregoing  
15 nucleotide sequence is reported in SEQ ID NO:186. Amino acids 15 to 27 are a  
predicted leader/signal sequence, with the predicted mature amino acid sequence  
beginning at amino acid 28, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
clone bm41\_7 should be approximately 1700 bp.

20 The nucleotide sequence disclosed herein for bm41\_7 was searched against the  
GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
FASTA search protocols. bm41\_7 demonstrated at least some similarity with sequences  
identified as AF047439 (Homo sapiens unknown mRNA, complete cds), H44519  
(yo74d10.r1 Homo sapiens cDNA clone 183667 5'), N29833 (yw93d10.s1 Homo sapiens  
cDNA clone 259795 3'), T23021 (Human gene signature HUMGS04750; standard; cDNA  
25 to mRNA), W58059 (zd22f10.r1 Soares fetal heart NbHH19W Homo sapiens cDNA  
clone), and Z78368 (H.sapiens mRNA, expressed sequence tag ICRFp507F18226, mRNA  
sequence). The predicted amino acid sequence disclosed herein for bm41\_7 was  
searched against the GenPept and GeneSeq amino acid sequence databases using the  
30 BLASTX search protocol. The predicted bm41\_7 protein demonstrated at least some  
similarity to the sequence identified as AF047439 (unknown [Homo sapiens]). Based  
upon sequence similarity, bm41\_7 proteins and each similar protein or peptide may  
share at least some activity. The TopPredII computer program predicts an additional  
potential trans-membrane domain within the bm41\_7 protein sequence centered  
35 around amino acid 252 of SEQ ID NO:186.

Clone "br342\_11"

A polynucleotide of the present invention has been identified as clone "br342\_11". br342\_11 was isolated from a human fetal kidney cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. br342\_11 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "br342\_11 protein").

The nucleotide sequence of br342\_11 as presently determined is reported in SEQ ID NO:187. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the br342\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:188. Amino acids 49 to 61 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 62, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone br342\_11 should be approximately 1400 bp.

The nucleotide sequence disclosed herein for br342\_11 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. br342\_11 demonstrated at least some similarity with sequences identified as Z69722 (Human DNA sequence from cosmid U212C1, between markers DXS366 and DXS87 on chromosome X), Z93019 Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 49C23; HTGS phase 1; Human DNA sequence from PAC 49C23 on chromosome X contains malate dehydrogenase pseudogene and STS), and Z95126 (Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 30P20; HTGS phase 1; Human DNA sequence from PAC 30P20 on chromosome Xq21.1-Xq21.3. Contains set pseudogene, ESTs and STS). The predicted amino acid sequence disclosed herein for br342\_11 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted br342\_11 protein demonstrated at least some similarity to sequences identified as D89049 (lectin-like oxidized LDL receptor [Bos taurus]) and R99586 (Low density lipoprotein receptor). Based upon sequence similarity, br342\_11 proteins and each similar protein or peptide may share at least some activity. The nucleotide sequence of br342\_11 indicates that it may contain one or more of the following repetitive elements: MER4A, MER4B.

Clone "ej258\_11"

A polynucleotide of the present invention has been identified as clone "ej258\_11". ej258\_11 was isolated from a human adult placenta cDNA library using

methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ej258\_11 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ej258\_11 protein").

The nucleotide sequence of ej258\_11 as presently determined is reported in SEQ ID NO:189. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ej258\_11 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:190.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ej258\_11 should be approximately 670 bp.

The nucleotide sequence disclosed herein for ej258\_11 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ej258\_11 demonstrated at least some similarity with sequences identified as AA217161 (mu86g11.r1 Soares mouse lymph node NbMLN Mus musculus cDNA clone 652484 5' similar to WP:F35H12.2 CE04511), AA330720 (EST34452 Embryo, 6 week I Homo sapiens cDNA 5' end), AJ000649 (Oryctolagus cuniculus unknown differentially expressed mRNA), U17432 (Bos taurus beta-mannosidase mRNA, complete cds), U91321 (Human chromosome 16p13 BAC clone CIT987SK-363E6, complete sequence), Z74031 (Caenorhabditis elegans cosmid F32D8), and Z99127 (Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 102G20; HTGS phase 1; Human DNA sequence from PAC 102G20 on chromosome 1q24-q25. Contains ESTS, STSs and a predicted CpG island). The predicted amino acid sequence disclosed herein for ej258\_11 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted ej258\_11 protein demonstrated at least some similarity to the sequence identified as U41540 (coded for by C. elegans cDNA yk42d12.5; coded for by C. elegans cDNA yk27e10.5; coded for by C. elegans cDNA cm08h6; coded for by C. elegans cDNA yk88e12.5). Based upon sequence similarity, ej258\_11 proteins and each similar protein or peptide may share at least some activity.

#### Clone "k232\_2x"

A polynucleotide of the present invention has been identified as clone "k232\_2x". A cDNA clone was first isolated from a murine adult bone marrow cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. This murine cDNA was then used to isolate k232\_2x, a full-length human

cDNA clone, including the entire coding sequence of a secreted protein (also referred to herein as "k232\_2x protein").

The nucleotide sequence of k232\_2x as presently determined is reported in SEQ ID NO:191. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the k232\_2x protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:192. Amino acids 4 to 16 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 17, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone k232\_2x should be approximately 555 bp.

The nucleotide sequence disclosed herein for k232\_2x was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. k232\_2x demonstrated at least some similarity with sequences identified as AA087828 (mn94b04.r1 Stratagene mouse lung 937302 Mus musculus cDNA clone 551695 5'), AA095731 (I5720.seq.F Fetal heart, Lambda ZAP Express Homo sapiens cDNA 5'), AA398859 (zt80e12.r1 Soares testis NHT Homo sapiens cDNA clone 728686 5'), N78829 (zb17a05.s1 Homo sapiens cDNA clone 302288 3'), T21965 (Human gene signature HUMGS03508), and W17346 (zb18c05.r1 Soares fetal lung NbHL19W Homo sapiens cDNA clone 302408 5'). Based upon sequence similarity, k232\_2x proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts two potential transmembrane domains within the k232\_2x protein sequence, one near the signal sequence and another near the C-terminus of SEQ ID NO:192.

#### Clone "lf307\_5"

A polynucleotide of the present invention has been identified as clone "lf307\_5". lf307\_5 was isolated from a human adult spinal cord cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. lf307\_5 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "lf307\_5 protein").

The nucleotide sequence of the 5' portion of lf307\_5 as presently determined is reported in SEQ ID NO:193. What applicants presently believe is the proper reading frame for the coding region is indicated in SEQ ID NO:194. The predicted amino acid sequence of the lf307\_5 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:194. Additional nucleotide sequence from the 3' portion of lf307\_5, including the polyA tail, is reported in SEQ ID NO:195.



The EcoRI/NotI restriction fragment obtainable from the deposit containing clone lf307\_5 should be approximately 1000 bp.

The nucleotide sequence disclosed herein for lf307\_5 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. lf307\_5 demonstrated at least some similarity with sequences identified as AA039895 (zk46a02.r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 485834 5') and AA513783 (nh89a05.r1 NCI\_CGAP\_Br1.1 Homo sapiens cDNA clone 965648). Based upon sequence similarity, lf307\_5 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the lf307\_5 protein sequence centered around amino acid 50 of SEQ ID NO:194.

#### Clone "lr204\_1"

A polynucleotide of the present invention has been identified as clone "lr204\_1". lr204\_1 was isolated from a human adult lymph node cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. lr204\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "lr204\_1 protein").

The nucleotide sequence of lr204\_1 as presently determined is reported in SEQ ID NO:196. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the lr204\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:197. Amino acids 29 to 41 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 42, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone lr204\_1 should be approximately 900 bp.

The nucleotide sequence disclosed herein for lr204\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. lr204\_1 demonstrated at least some similarity with sequences identified as N47763 (yy55e07.r1 Homo sapiens cDNA clone 277476 5') and N56875 (yy55e07.s1 Homo sapiens cDNA clone 277476 3' similar to SW:CYTO\_BOVIN P01035 CYSTATIN, COLOSTRUM). The predicted amino acid sequence disclosed herein for lr204\_1 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted lr204\_1 protein demonstrated at least some similarity to sequences identified as M27891 (cystatin C [Homo sapiens]), P94392

(Sequence of complete recombinant cystatin C in E.coli), R43323 (Cystatin polypeptide), and X62412 (cystatin [unidentified]). Based upon sequence similarity, lr204\_1 proteins and each similar protein or peptide may share at least some activity. The predicted lr204\_1 protein contains a sequence highly conserved in all cystatins (QIVAG in human cystatin C, QIVKG in the predicted lr204\_1 protein). Cystatins are inhibitors of papain-like cysteine proteinases such as cathepsins. Cystatin C belongs to family 2 of the cystatin superfamily. The family 2 cystatins are secreted proteins of about 120 amino acids. All cystatins have important roles in processes involving cysteine proteinase activity like bone resorption. They are also implicated in a variety of diseases (e.g. sepsis, cancer metastasis, rheumatoid arthritis etc.) since they regulate potentially harmful proteinase activity. The predicted lr204\_1 protein appears to be a novel cystatin C related inhibitor of cysteine proteinases. The TopPredII computer program predicts a potential transmembrane domain within the lr204\_1 protein sequence near amino acid 42 of SEQ ID NO:197.

15

#### Clone "as20\_2"

A polynucleotide of the present invention has been identified as clone "as20\_2". as20\_2 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. as20\_2 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "as20\_2 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of as20\_2 as presently determined is reported in SEQ ID NO:205. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the as20\_2 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:206. Amino acids 17 to 29 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 30, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone as20\_2 should be approximately 2100 bp.

The nucleotide sequence disclosed herein for as20\_2 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. as20\_2 demonstrated at least some similarity with sequences identified as AA192606 (zq01g04.s1 Stratagene muscle 937209 Homo sapiens cDNA clone 628470 3'), AF000657 (Arabidopsis thaliana BAC F19G10, complete sequence), T91778 (yd52c10.s1 Homo sapiens cDNA clone 111858 3' similar to contains Alu repetitive element), and W26193 (22b2 Human retina cDNA randomly primed

sublibrary Homo sapiens cDNA). Based upon sequence similarity, as20\_2 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the as20\_2 protein sequence centered around amino acid 97 of SEQ ID NO:206.

#### Clone "bf227\_8"

A polynucleotide of the present invention has been identified as clone "bf227\_8". bf227\_8 was isolated from a human fetal brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bf227\_8 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bf227\_8 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of bf227\_8 as presently determined is reported in SEQ ID NO:207. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bf227\_8 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:208.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bf227\_8 should be approximately 1400 bp.

The nucleotide sequence disclosed herein for bf227\_8 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bf227\_8 demonstrated at least some similarity with sequences identified as AA452345 (zx15c09.r1 Soares total fetus Nb2HF8 9w Homo sapiens cDNA clone 786544 5'), N20474 (yx39d10.s1 Homo sapiens cDNA clone 264115 3'), N79685 (yz82a08.r1 Homo sapiens cDNA clone 289526 5'), and W73775 (zd50d09.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 344081 3'). The predicted amino acid sequence disclosed herein for bf227\_8 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bf227\_8 protein demonstrated at least some similarity to sequences identified as U11768 (coat protein [Grapevine fanleaf virus]). Based upon sequence similarity, bf227\_8 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the bf227\_8 protein sequence centered around amino acid 22 of SEQ ID NO:208.

#### Clone "bh157\_7"

A polynucleotide of the present invention has been identified as clone "bh157\_7". bh157\_7 was isolated from a human adult ovary cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No.

5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. bh157\_7 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "bh157\_7 protein") and a poly(A) tail at its 3' end.

5 The nucleotide sequence of bh157\_7 as presently determined is reported in SEQ ID NO:209. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the bh157\_7 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:210.

10 The EcoRI/NotI restriction fragment obtainable from the deposit containing clone bh157\_7 should be approximately 1250 bp.

The nucleotide sequence disclosed herein for bh157\_7 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. bh157\_7 demonstrated at least some similarity with sequences identified as AA312435 (EST183106 Jurkat T-cells VI Homo sapiens cDNA 5' end) and T62753 (yc70g05.r1 Homo sapiens cDNA clone 86072 5'). The predicted amino acid sequence disclosed herein for bh157\_7 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted bh157\_7 protein demonstrated at least some similarity to sequences identified as X84037 (E-selectin ligand-1 [Mus musculus]). Based upon sequence similarity, 20 bh157\_7 proteins and each similar protein or peptide may share at least some activity.

#### Clone "cg426\_8"

A polynucleotide of the present invention has been identified as clone "cg426\_8". cg426\_8 was isolated from a human adult testes cDNA library using 25 methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. cg426\_8 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "cg426\_8 protein") and a poly(A) tail at its 3' end.

30 The nucleotide sequence of cg426\_8 as presently determined is reported in SEQ ID NO:211. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the cg426\_8 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:212. Amino acids 4 to 16 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at 35 amino acid 17, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone cg426\_8 should be approximately 2500 bp.

The nucleotide sequence disclosed herein for cg426\_8 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. cg426\_8 demonstrated at least some similarity with sequences identified as AA523415 (ng30a08.s1 NCI\_CGAP\_Co3 Homo sapiens cDNA clone  
 5 936278 similar to contains element MER22 repetitive element), N58694 (yv64f11.r1 Homo sapiens cDNA clone 247533 5'), and W78817 (zh51c03.r1 Soares fetal liver spleen 1NFLS S1 Homo sapiens cDNA clone 415588 5' similar to contains MER1.t3 MER1 repetitive element). Based upon sequence similarity, cg426\_8 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program  
 10 predicts a potential transmembrane domain within the cg426\_8 protein sequence centered around amino acid 55 of SEQ ID NO:212. The nucleotide sequence of cg426\_8 may contain a MER repetitive element.

#### Clone "ck48\_12"

15 A polynucleotide of the present invention has been identified as clone "ck48\_12". ck48\_12 was isolated from a human adult testes cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ck48\_12  
 20 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ck48\_12 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of ck48\_12 as presently determined is reported in SEQ ID NO:213. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ck48\_12 protein corresponding to the foregoing  
 25 nucleotide sequence is reported in SEQ ID NO:214. Amino acids 119 to 131 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 132, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ck48\_12 should be approximately 1350 bp.

30 The nucleotide sequence disclosed herein for ck48\_12 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ck48\_12 demonstrated at least some similarity with sequences identified as AA064481 (ml50a08.r1 Stratagene mouse testis (#937308) Mus musculus cDNA clone 515414 5') and AA397716 (zt87f10.r1 Soares testis NHT Homo sapiens  
 35 cDNA clone 729355 5'). Based upon sequence similarity, ck48\_12 proteins and each similar protein or peptide may share at least some activity. The ck48\_12 protein sequence may have a transmembrane domain at the carboxyl terminus of SEQ ID NO:214.

Clone "co1000\_1"

A polynucleotide of the present invention has been identified as clone "co1000\_1". co1000\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. co1000\_1 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "co1000\_1 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of co1000\_1 as presently determined is reported in SEQ ID NO:215. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the co1000\_1 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:216. Amino acids 14 to 26 are a predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 27, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone co1000\_1 should be approximately 1600 bp.

The nucleotide sequence disclosed herein for co1000\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. co1000\_1 demonstrated at least some similarity with sequences identified as M77867 (EST01451 Homo sapiens cDNA clone HFBCA06 similar to Alu repetitive element). Based upon sequence similarity, co1000\_1 proteins and each similar protein or peptide may share at least some activity. The nucleotide sequence of co1000\_1 indicates that it may contain Alu and Mer4 repetitive elements.

Clone "ct489\_14"

A polynucleotide of the present invention has been identified as clone "ct489\_14". ct489\_14 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. ct489\_14 is a full-length clone, including the entire coding sequence of a secreted protein (also referred to herein as "ct489\_14 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of ct489\_14 as presently determined is reported in SEQ ID NO:217. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the ct489\_14 protein corresponding to the foregoing nucleotide sequence is reported in SEQ ID NO:218. Amino acids 12 to 24 are a

predicted leader/signal sequence, with the predicted mature amino acid sequence beginning at amino acid 25, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone ct489\_14 should be approximately 1900 bp.

5        The nucleotide sequence disclosed herein for ct489\_14 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. ct489\_14 demonstrated at least some similarity with sequences identified as H21179 (yn66d09.s1 Homo sapiens cDNA clone 173393 3'), N99345 (IMAGE:59425 Homo sapiens cDNA clone 59425), and R89669 (ym97f05.r1  
10 Homo sapiens cDNA clone 166881 5'). The predicted amino acid sequence disclosed herein for ct489\_14 was searched against the GenPept and GeneSeq amino acid sequence databases using the BLASTX search protocol. The predicted ct489\_14 protein demonstrated at least some similarity to sequences identified as U23803 (heterogeneous ribonucleoprotein A0 [Homo sapiens]). Based upon sequence similarity, ct489\_14  
15 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the ct489\_14 protein sequence centered around amino acid 280 of SEQ ID NO:217.

#### Clone "df821\_1"

20        A polynucleotide of the present invention has been identified as clone "df821\_1". df821\_1 was isolated from a human adult brain cDNA library using methods which are selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was identified as encoding a secreted or transmembrane protein on the basis of computer analysis of the amino acid sequence of the encoded protein. df821\_1 is a full-length  
25 clone, including the entire coding sequence of a secreted protein (also referred to herein as "df821\_1 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of df821\_1 as presently determined is reported in SEQ ID NO:219. What applicants presently believe to be the proper reading frame and the predicted amino acid sequence of the df821\_1 protein corresponding to the foregoing  
30 nucleotide sequence is reported in SEQ ID NO:220.

The EcoRI/NotI restriction fragment obtainable from the deposit containing clone df821\_1 should be approximately 800 bp.

35        The nucleotide sequence disclosed herein for df821\_1 was searched against the GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and FASTA search protocols. df821\_1 demonstrated at least some similarity with sequences identified as AA311729 (EST182669 Jurkat T-cells VI Homo sapiens cDNA 5' end similar to similar to hypothetical protein pIL2), M17412 (Rat growth and transformation- dependent mRNA, 3' end), W01700 (za37a03.r1 Soares fetal liver spleen

1NFLS Homo sapiens cDNA clone 294700 5' similar to PIR A26882 A26882 pIL2  
 hypothetical protein - rat), W44481 (zc28g12.r1 Soares senescent fibroblasts NbHSF  
 Homo sapiens cDNA clone 323686 5' similar to PIR:A26882 A26882 pIL2 hypothetical  
 protein - rat), and W93991 (zd98b04.s1 Soares fetal heart NbHH19W Homo sapiens  
 5 cDNA clone 357487 3' similar to PIR A26882 A26882 pIL2 hypothetical protein - rat).  
 The predicted amino acid sequence disclosed herein for df821\_1 was searched against  
 the GenPept and GeneSeq amino acid sequence databases using the BLASTX search  
 protocol. The predicted df821\_1 protein demonstrated at least some similarity to  
 sequences identified as M17412 (growth and transformation dependent protein [Rattus  
 10 norvegicus]). Based upon sequence similarity, df821\_1 proteins and each similar  
 protein or peptide may share at least some activity. The TopPredII computer program  
 predicts a potential transmembrane domain within the df821\_1 protein sequence  
 centered around amino acid 110 of SEQ ID NO:220.

15 Clone "dy41\_2"

A polynucleotide of the present invention has been identified as clone "dy41\_2".  
 dy41\_2 was isolated from a human adult brain cDNA library using methods which are  
 selective for cDNAs encoding secreted proteins (see U.S. Pat. No. 5,536,637), or was  
 identified as encoding a secreted or transmembrane protein on the basis of computer  
 20 analysis of the amino acid sequence of the encoded protein. dy41\_2 is a full-length  
 clone, including the entire coding sequence of a secreted protein (also referred to herein  
 as "dy41\_2 protein") and a poly(A) tail at its 3' end.

The nucleotide sequence of dy41\_2 as presently determined is reported in SEQ  
 ID NO:221. What applicants presently believe to be the proper reading frame and the  
 25 predicted amino acid sequence of the dy41\_2 protein corresponding to the foregoing  
 nucleotide sequence is reported in SEQ ID NO:222. Amino acids 32 to 44 are a  
 predicted leader/signal sequence, with the predicted mature amino acid sequence  
 beginning at amino acid 45, or are within a transmembrane domain.

The EcoRI/NotI restriction fragment obtainable from the deposit containing  
 30 clone dy41\_2 should be approximately 1200 bp.

The nucleotide sequence disclosed herein for dy41\_2 was searched against the  
 GenBank and GeneSeq nucleotide sequence databases using BLASTN/BLASTX and  
 FASTA search protocols. dy41\_2 demonstrated at least some similarity with sequences  
 identified as H02323 (yj40f04.s1 Homo sapiens cDNA clone 151231 3'). The predicted  
 35 amino acid sequence disclosed herein for dy41\_2 was searched against the GenPept and  
 GeneSeq amino acid sequence databases using the BLASTX search protocol. The  
 predicted dy41\_2 protein demonstrated at least some similarity to sequences identified  
 as D89050 (lectin-like oxidized LDL receptor [Homo sapiens]). Based upon sequence



similarity, dy41\_2 proteins and each similar protein or peptide may share at least some activity. The TopPredII computer program predicts a potential transmembrane domain within the dy41\_2 protein sequence, from amino acid 40 to amino acid 60 of SEQ ID NO:222.

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#### Deposit of Clones

Clones bh389\_11, bk112\_15, bk200\_13, di386\_3, em397\_2, fh170\_7, fn53\_4, fq505\_4, fw13\_9, and gg619\_2 were deposited on June 10, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98451, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones cl181\_3, cr1044\_1, cz251\_1, dd12\_7, fn191\_3, gm196\_4, gn114\_1, hj968\_2, hk10\_3, and hm236\_1 were deposited on June 12, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98456, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones do15\_4, dx290\_1, ek390\_4, er471\_7, fs40\_3, ga63\_6, gm335\_4, hy370\_9, ie47\_4, and s195\_10 were deposited on June 19, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98468, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones bf228\_14, bg249\_1, bv286\_1, co36\_1, cp116\_1, cw1195\_2, fh13\_10, gc57\_4, h1165\_3 and hb752\_1 were deposited on July 2, 1997 with the American Type Culture Collection as an original deposit under the Budapest Treaty and were given the accession number ATCC 98482, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the

deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b).

Clones bi127\_5, bl194\_2, cc130\_1, ch582\_1, cq294\_14, dd454\_1, du157\_9, du372\_1, ej90\_5, and ic2\_6 were deposited on August 5, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98501, from which each clone comprising a particular polynucleotide is obtainable. Clone du157\_12 is an additional isolate of clone du157\_9 and was deposited on April 7, 1998 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and was given the accession number ATCC 98724, from which the du157\_12 clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones bn97\_1, bn268\_11, cb96\_10, cb213\_11, cj457\_4, cz653\_11, dx138\_4, and ij167\_5 were deposited on September 4, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98535, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones bd107\_16 and bm41\_7 were deposited on September 25, 1998 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number ATCC 98898, from which each clone comprising a particular polynucleotide is obtainable. Clones bd107\_13, bm41\_3, br342\_11, ej258\_11, k232\_2x, lf307\_5, and lr204\_1 were deposited on October 2, 1997 with the American Type Culture Collection as an original deposit under the Budapest Treaty and were given the accession number ATCC 98551, from which each clone comprising a particular polynucleotide is obtainable. Clones bd107\_13 and bm41\_3 are additional isolates of clones bd107\_16 and bm41\_7, respectively. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

Clones as20\_2, bf227\_8, bh157\_7, cg426\_8, ck48\_12, co1000\_1, ct489\_14, df821\_1 and dy41\_2 were deposited on November 7, 1997 with the American Type Culture Collection (10801 University Boulevard, Manassas, Virginia 20110-2209 U.S.A.) as an original deposit under the Budapest Treaty and were given the accession number  
 5 ATCC 98580, from which each clone comprising a particular polynucleotide is obtainable. All restrictions on the availability to the public of the deposited material will be irrevocably removed upon the granting of the patent, except for the requirements specified in 37 C.F.R. § 1.808(b), and the term of the deposit will comply with 37 C.F.R. § 1.806.

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Each clone has been transfected into separate bacterial cells (*E. coli*) in this composite deposit. Each clone can be removed from the vector in which it was deposited by performing an EcoRI/NotI digestion (5' site, EcoRI; 3' site, NotI) to produce the appropriate fragment for such clone. Each clone was deposited in either  
 15 the pED6 or pNOTs vector depicted in Figures 1A and 1B, respectively. The pED6dpc2 vector ("pED6") was derived from pED6dpc1 by insertion of a new polylinker to facilitate cDNA cloning (Kaufman *et al.*, 1991, *Nucleic Acids Res.* 19: 4485-4490); the pNOTs vector was derived from pMT2 (Kaufman *et al.*, 1989, *Mol. Cell. Biol.* 9: 946-958) by deletion of the DHFR sequences, insertion of a new polylinker, and insertion of the  
 20 M13 origin of replication in the ClaI site. In some instances, the deposited clone can become "flipped" (i.e., in the reverse orientation) in the deposited isolate. In such instances, the cDNA insert can still be isolated by digestion with EcoRI and NotI. However, NotI will then produce the 5' site and EcoRI will produce the 3' site for placement of the cDNA in proper orientation for expression in a suitable vector. The  
 25 cDNA may also be expressed from the vectors in which they were deposited.

Bacterial cells containing a particular clone can be obtained from the composite deposit as follows:

An oligonucleotide probe or probes should be designed to the sequence that is known for that particular clone. This sequence can be derived from the sequences  
 30 provided herein, or from a combination of those sequences. The sequence of an oligonucleotide probe that was used to isolate or to sequence each full-length clone is identified below, and should be most reliable in isolating the clone of interest.

Clone	Probe Sequence
35 bh389_11	SEQ ID NO:24
bk112_15	SEQ ID NO:25
bk200_13	SEQ ID NO:26
di386_3	SEQ ID NO:27

	em397_2	SEQ ID NO:28
	fh170_7	SEQ ID NO:29
	fn53_4	SEQ ID NO:30
	fq505_4	SEQ ID NO:31
5	fw13_9	SEQ ID NO:32
	gg619_2	SEQ ID NO:33
	cl181_3	SEQ ID NO:55
	cr1044_1	SEQ ID NO:56
	cz251_1	SEQ ID NO:57
10	dd12_7	SEQ ID NO:58
	fn191_3	SEQ ID NO:59
	gm196_4	SEQ ID NO:60
	gn114_1	SEQ ID NO:61
	hj968_2	SEQ ID NO:62
15	hk10_3	SEQ ID NO:63
	hm236_1	SEQ ID NO:64
	do15_4	SEQ ID NO:87
	dx290_1	SEQ ID NO:88
	ek390_4	SEQ ID NO:89
20	er471_7	SEQ ID NO:90
	fs40_3	SEQ ID NO:91
	ga63_6	SEQ ID NO:92
	gm335_4	SEQ ID NO:93
	hy370_9	SEQ ID NO:94
25	ie47_4	SEQ ID NO:95
	s195_10	SEQ ID NO:96
	bf228_14	SEQ ID NO:119
	bg249_1	SEQ ID NO:120
	bv286_1	SEQ ID NO:121
30	co36_1	SEQ ID NO:122
	cp116_1	SEQ ID NO:123
	cw1195_2	SEQ ID NO:124
	fh13_10	SEQ ID NO:125
	gc57_4	SEQ ID NO:126
35	h1165_3	SEQ ID NO:127
	hb752_1	SEQ ID NO:128
	bi127_5	SEQ ID NO:149
	bl194_2	SEQ ID NO:150

	cc130_1	SEQ ID NO:151
	ch582_1	SEQ ID NO:152
	cq294_14	SEQ ID NO:153
	dd454_1	SEQ ID NO:154
5	du157_12	SEQ ID NO:155
	du372_1	SEQ ID NO:156
	ej90_5	SEQ ID NO:157
	ic2_6	SEQ ID NO:158
	bn97_1	SEQ ID NO:175
10	bn268_11	SEQ ID NO:176
	cb96_10	SEQ ID NO:177
	cb213_11	SEQ ID NO:178
	cj457_4	SEQ ID NO:179
	cz653_11	SEQ ID NO:180
15	dx138_4	SEQ ID NO:181
	ij167_5	SEQ ID NO:182
	bd107_16	SEQ ID NO:198
	bm41_7	SEQ ID NO:199
	br342_11	SEQ ID NO:200
20	ej258_11	SEQ ID NO:201
	k232_2x	SEQ ID NO:202
	lf307_5	SEQ ID NO:203
	lr204_1	SEQ ID NO:204
	as20_2	SEQ ID NO:223
25	bf227_8	SEQ ID NO:224
	bh157_7	SEQ ID NO:225
	cg426_8	SEQ ID NO:226
	ck48_12	SEQ ID NO:227
	co1000_1	SEQ ID NO:228
30	ct489_14	SEQ ID NO:229
	df821_1	SEQ ID NO:230
	dy41_2	SEQ ID NO:231

In the sequences listed above which include an N at position 2, that position is occupied in preferred probes/primers by a biotinylated phosphoramidite residue rather than a nucleotide (such as , for example, that produced by use of biotin phosphoramidite (1-dimethoxytrityloxy-2-(N-biotinyl-4-aminobutyl)-propyl-3-O-(2-cyanoethyl)-(N,N-diisopropyl)-phosphoramidite) (Glen Research, cat. no. 10-1953)).

The design of the oligonucleotide probe should preferably follow these parameters:

- (a) It should be designed to an area of the sequence which has the fewest ambiguous bases ("N's"), if any;
- 5 (b) It should be designed to have a  $T_m$  of approx. 80 ° C (assuming 2° for each A or T and 4 degrees for each G or C).

The oligonucleotide should preferably be labeled with  $\gamma$ - $^{32}\text{P}$  ATP (specific activity 6000 Ci/mmole) and T4 polynucleotide kinase using commonly employed techniques for labeling oligonucleotides. Other labeling techniques can also be used. Unincorporated  
 10 label should preferably be removed by gel filtration chromatography or other established methods. The amount of radioactivity incorporated into the probe should be quantitated by measurement in a scintillation counter. Preferably, specific activity of the resulting probe should be approximately  $4\text{e}+6$  dpm/pmole.

The bacterial culture containing the pool of full-length clones should preferably  
 15 be thawed and 100  $\mu\text{l}$  of the stock used to inoculate a sterile culture flask containing 25 ml of sterile L-broth containing ampicillin at 100  $\mu\text{g}/\text{ml}$ . The culture should preferably be grown to saturation at 37°C, and the saturated culture should preferably be diluted in fresh L-broth. Aliquots of these dilutions should preferably be plated to determine the dilution and volume which will yield approximately 5000 distinct and well-  
 20 separated colonies on solid bacteriological media containing L-broth containing ampicillin at 100  $\mu\text{g}/\text{ml}$  and agar at 1.5% in a 150 mm petri dish when grown overnight at 37°C. Other known methods of obtaining distinct, well-separated colonies can also be employed.

Standard colony hybridization procedures should then be used to transfer the  
 25 colonies to nitrocellulose filters and lyse, denature and bake them.

The filter is then preferably incubated at 65°C for 1 hour with gentle agitation in 6X SSC (20X stock is 175.3 g NaCl/liter, 88.2 g Na citrate/liter, adjusted to pH 7.0 with NaOH) containing 0.5% SDS, 100  $\mu\text{g}/\text{ml}$  of yeast RNA, and 10 mM EDTA (approximately 10 mL per 150 mm filter). Preferably, the probe is then added to the  
 30 hybridization mix at a concentration greater than or equal to  $1\text{e}+6$  dpm/mL. The filter is then preferably incubated at 65°C with gentle agitation overnight. The filter is then preferably washed in 500 mL of 2X SSC/0.5% SDS at room temperature without agitation, preferably followed by 500 mL of 2X SSC/0.1% SDS at room temperature with gentle shaking for 15 minutes. A third wash with 0.1X SSC/0.5% SDS at 65°C for  
 35 30 minutes to 1 hour is optional. The filter is then preferably dried and subjected to autoradiography for sufficient time to visualize the positives on the X-ray film. Other known hybridization methods can also be employed.

The positive colonies are picked, grown in culture, and plasmid DNA isolated using standard procedures. The clones can then be verified by restriction analysis, hybridization analysis, or DNA sequencing.

Fragments of the proteins of the present invention which are capable of exhibiting biological activity are also encompassed by the present invention. Fragments of the protein may be in linear form or they may be cyclized using known methods, for example, as described in H.U. Saragovi, *et al.*, Bio/Technology 10, 773-778 (1992) and in R.S. McDowell, *et al.*, J. Amer. Chem. Soc. 114, 9245-9253 (1992), both of which are incorporated herein by reference. Such fragments may be fused to carrier molecules such as immunoglobulins for many purposes, including increasing the valency of protein binding sites. For example, fragments of the protein may be fused through "linker" sequences to the Fc portion of an immunoglobulin. For a bivalent form of the protein, such a fusion could be to the Fc portion of an IgG molecule. Other immunoglobulin isotypes may also be used to generate such fusions. For example, a protein - IgM fusion would generate a decavalent form of the protein of the invention.

The present invention also provides both full-length and mature forms of the disclosed proteins. The full-length form of the such proteins is identified in the sequence listing by translation of the nucleotide sequence of each disclosed clone. The mature form(s) of such protein may be obtained by expression of the disclosed full-length polynucleotide (preferably those deposited with ATCC) in a suitable mammalian cell or other host cell. The sequence(s) of the mature form(s) of the protein may also be determinable from the amino acid sequence of the full-length form.

The present invention also provides genes corresponding to the polynucleotide sequences disclosed herein. "Corresponding genes" are the regions of the genome that are transcribed to produce the mRNAs from which cDNA polynucleotide sequences are derived and may include contiguous regions of the genome necessary for the regulated expression of such genes. Corresponding genes may therefore include but are not limited to coding sequences, 5' and 3' untranslated regions, alternatively spliced exons, introns, promoters, enhancers, and silencer or suppressor elements. The corresponding genes can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include the preparation of probes or primers from the disclosed sequence information for identification and/or amplification of genes in appropriate genomic libraries or other sources of genomic materials. An "isolated gene" is a gene that has been separated from the adjacent coding sequences, if any, present in the genome of the organism from which the gene was isolated.

Organisms that have enhanced, reduced, or modified expression of the gene(s) corresponding to the polynucleotide sequences disclosed herein are provided. The

desired change in gene expression can be achieved through the use of antisense polynucleotides or ribozymes that bind and/or cleave the mRNA transcribed from the gene (Albert and Morris, 1994, *Trends Pharmacol. Sci.* **15**(7): 250-254; Lavarosky *et al.*, 1997, *Biochem. Mol. Med.* **62**(1): 11-22; and Hampel, 1998, *Prog. Nucleic Acid Res. Mol. Biol.* **58**: 1- 39; all of which are incorporated by reference herein). Transgenic animals that have multiple copies of the gene(s) corresponding to the polynucleotide sequences disclosed herein, preferably produced by transformation of cells with genetic constructs that are stably maintained within the transformed cells and their progeny, are provided. Transgenic animals that have modified genetic control regions that increase or reduce gene expression levels, or that change temporal or spatial patterns of gene expression, are also provided (see European Patent No. 0 649 464 B1, incorporated by reference herein). In addition, organisms are provided in which the gene(s) corresponding to the polynucleotide sequences disclosed herein have been partially or completely inactivated, through insertion of extraneous sequences into the corresponding gene(s) or through deletion of all or part of the corresponding gene(s). Partial or complete gene inactivation can be accomplished through insertion, preferably followed by imprecise excision, of transposable elements (Plasterk, 1992, *Bioessays* **14**(9): 629-633; Zwaal *et al.*, 1993, *Proc. Natl. Acad. Sci. USA* **90**(16): 7431-7435; Clark *et al.*, 1994, *Proc. Natl. Acad. Sci. USA* **91**(2): 719-722; all of which are incorporated by reference herein), or through homologous recombination, preferably detected by positive/negative genetic selection strategies (Mansour *et al.*, 1988, *Nature* **336**: 348-352; U.S. Patent Nos. 5,464,764; 5,487,992; 5,627,059; 5,631,153; 5,614, 396; 5,616,491; and 5,679,523; all of which are incorporated by reference herein). These organisms with altered gene expression are preferably eukaryotes and more preferably are mammals. Such organisms are useful for the development of non-human models for the study of disorders involving the corresponding gene(s), and for the development of assay systems for the identification of molecules that interact with the protein product(s) of the corresponding gene(s).

Where the protein of the present invention is membrane-bound (e.g., is a receptor), the present invention also provides for soluble forms of such protein. In such forms part or all of the intracellular and transmembrane domains of the protein are deleted such that the protein is fully secreted from the cell in which it is expressed. The intracellular and transmembrane domains of proteins of the invention can be identified in accordance with known techniques for determination of such domains from sequence information.

Proteins and protein fragments of the present invention include proteins with amino acid sequence lengths that are at least 25%(more preferably at least 50%, and most preferably at least 75%) of the length of a disclosed protein and have at least 60%



sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with that disclosed protein, where sequence identity is determined by comparing the amino acid sequences of the proteins when aligned so as to maximize overlap and identity while minimizing sequence gaps. Also included in the present invention are proteins and protein fragments that contain a segment preferably comprising 8 or more (more preferably 20 or more, most preferably 30 or more) contiguous amino acids that shares at least 75% sequence identity (more preferably, at least 85% identity; most preferably at least 95% identity) with any such segment of any of the disclosed proteins.

Species homologues of the disclosed polynucleotides and proteins are also provided by the present invention. As used herein, a "species homologue" is a protein or polynucleotide with a different species of origin from that of a given protein or polynucleotide, but with significant sequence similarity to the given protein or polynucleotide. Preferably, polynucleotide species homologues have at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% identity) with the given polynucleotide, and protein species homologues have at least 30% sequence identity (more preferably, at least 45% identity; most preferably at least 60% identity) with the given protein, where sequence identity is determined by comparing the nucleotide sequences of the polynucleotides or the amino acid sequences of the proteins when aligned so as to maximize overlap and identity while minimizing sequence gaps. Species homologues may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source from the desired species. Preferably, species homologues are those isolated from mammalian species. Most preferably, species homologues are those isolated from certain mammalian species such as, for example, *Pan troglodytes*, *Gorilla gorilla*, *Pongo pygmaeus*, *Hylobates concolor*, *Macaca mulatta*, *Papio papio*, *Papio hamadryas*, *Cercopithecus aethiops*, *Cebus capucinus*, *Aotus trivirgatus*, *Sanguinus oedipus*, *Microcebus murinus*, *Mus musculus*, *Rattus norvegicus*, *Cricetulus griseus*, *Felis catus*, *Mustela vison*, *Canis familiaris*, *Oryctolagus cuniculus*, *Bos taurus*, *Ovis aries*, *Sus scrofa*, and *Equus caballus*, for which genetic maps have been created allowing the identification of syntenic relationships between the genomic organization of genes in one species and the genomic organization of the related genes in another species (O'Brien and Seuánez, 1988, *Ann. Rev. Genet.* **22**: 323-351; O'Brien *et al.*, 1993, *Nature Genetics* **3**:103-112; Johansson *et al.*, 1995, *Genomics* **25**: 682-690; Lyons *et al.*, 1997, *Nature Genetics* **15**: 47-56; O'Brien *et al.*, 1997, *Trends in Genetics* **13**(10): 393-399; Carver and Stubbs, 1997, *Genome Research* **7**:1123-1137; all of which are incorporated by reference herein).

The invention also encompasses allelic variants of the disclosed polynucleotides or proteins; that is, naturally-occurring alternative forms of the isolated polynucleotides which also encode proteins which are identical or have significantly similar sequences to those encoded by the disclosed polynucleotides. Preferably, allelic variants have at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% identity) with the given polynucleotide, where sequence identity is determined by comparing the nucleotide sequences of the polynucleotides when aligned so as to maximize overlap and identity while minimizing sequence gaps. Allelic variants may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source from individuals of the appropriate species.

The invention also includes polynucleotides with sequences complementary to those of the polynucleotides disclosed herein.

The present invention also includes polynucleotides that hybridize under reduced stringency conditions, more preferably stringent conditions, and most preferably highly stringent conditions, to polynucleotides described herein. Examples of stringency conditions are shown in the table below: highly stringent conditions are those that are at least as stringent as, for example, conditions A-F; stringent conditions are at least as stringent as, for example, conditions G-L; and reduced stringency conditions are at least as stringent as, for example, conditions M-R.

Stringency Condition	Polynucleotide Hybrid	Hybrid Length (bp) <sup>‡</sup>	Hybridization Temperature and Buffer <sup>†</sup>	Wash Temperature and Buffer <sup>†</sup>
A	DNA:DNA	≥ 50	65°C; 1xSSC -or- 42°C; 1xSSC, 50% formamide	65°C; 0.3xSSC
B	DNA:DNA	<50	T <sub>B</sub> *; 1xSSC	T <sub>B</sub> *; 1xSSC
C	DNA:RNA	≤ 50	67°C; 1xSSC -or- 45°C; 1xSSC, 50% formamide	67°C; 0.3xSSC
D	DNA:RNA	<50	T <sub>D</sub> *; 1xSSC	T <sub>D</sub> *; 1xSSC
E	RNA:RNA	≥ 50	70°C; 1xSSC -or- 50°C; 1xSSC, 50% formamide	70°C; 0.3xSSC
F	RNA:RNA	<50	T <sub>F</sub> *; 1xSSC	T <sub>F</sub> *; 1xSSC
G	DNA:DNA	≥ 50	65°C; 4xSSC -or- 42°C; 4xSSC, 50% formamide	65°C; 1xSSC
H	DNA:DNA	<50	T <sub>H</sub> *; 4xSSC	T <sub>H</sub> *; 4xSSC
I	DNA:RNA	≥ 50	67°C; 4xSSC -or- 45°C; 4xSSC, 50% formamide	67°C; 1xSSC
J	DNA:RNA	<50	T <sub>J</sub> *; 4xSSC	T <sub>J</sub> *; 4xSSC
K	RNA:RNA	≥ 50	70°C; 4xSSC -or- 50°C; 4xSSC, 50% formamide	67°C; 1xSSC
L	RNA:RNA	<50	T <sub>L</sub> *; 2xSSC	T <sub>L</sub> *; 2xSSC
M	DNA:DNA	≥ 50	50°C; 4xSSC -or- 40°C; 6xSSC, 50% formamide	50°C; 2xSSC
N	DNA:DNA	<50	T <sub>N</sub> *; 6xSSC	T <sub>N</sub> *; 6xSSC
O	DNA:RNA	≥ 50	55°C; 4xSSC -or- 42°C; 6xSSC, 50% formamide	55°C; 2xSSC
P	DNA:RNA	<50	T <sub>P</sub> *; 6xSSC	T <sub>P</sub> *; 6xSSC
Q	RNA:RNA	≥ 50	60°C; 4xSSC -or-	60°C; 2xSSC

			45°C; 6xSSC, 50% formamide	
R	RNA:RNA	<50	T <sub>R</sub> *; 4xSSC	T <sub>R</sub> *; 4xSSC

†: The hybrid length is that anticipated for the hybridized region(s) of the hybridizing polynucleotides. When hybridizing a polynucleotide to a target polynucleotide of unknown sequence, the hybrid length is assumed to be that of the hybridizing polynucleotide. When polynucleotides of known sequence are hybridized, the hybrid length can be determined by aligning the sequences of the polynucleotides and identifying the region or regions of optimal sequence complementarity.

†: SSPE (1xSSPE is 0.15M NaCl, 10mM NaH<sub>2</sub>PO<sub>4</sub>, and 1.25mM EDTA, pH 7.4) can be substituted for SSC (1xSSC is 0.15M NaCl and 15mM sodium citrate) in the hybridization and wash buffers; washes are performed for 15 minutes after hybridization is complete.

\*T<sub>B</sub> - T<sub>R</sub>: The hybridization temperature for hybrids anticipated to be less than 50 base pairs in length should be 5-10°C less than the melting temperature (T<sub>m</sub>) of the hybrid, where T<sub>m</sub> is determined according to the following equations. For hybrids less than 18 base pairs in length, T<sub>m</sub>(°C) = 2(# of A + T bases) + 4(# of G + C bases). For hybrids between 18 and 49 base pairs in length, T<sub>m</sub>(°C) = 81.5 + 16.6(log<sub>10</sub>[Na<sup>+</sup>]) + 0.41(%G+C) - (600/N), where N is the number of bases in the hybrid, and [Na<sup>+</sup>] is the concentration of sodium ions in the hybridization buffer ([Na<sup>+</sup>] for 1xSSC = 0.165 M).

Additional examples of stringency conditions for polynucleotide hybridization are provided in Sambrook, J., E.F. Fritsch, and T. Maniatis, 1989, *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, chapters 9 and 11, and *Current Protocols in Molecular Biology*, 1995, F.M. Ausubel et al., eds., John Wiley & Sons, Inc., sections 2.10 and 6.3-6.4, incorporated herein by reference.

Preferably, each such hybridizing polynucleotide has a length that is at least 25%(more preferably at least 50%, and most preferably at least 75%) of the length of the polynucleotide of the present invention to which it hybridizes, and has at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with the polynucleotide of the present invention to which it hybridizes, where sequence identity is determined by comparing the sequences of the hybridizing polynucleotides when aligned so as to maximize overlap and identity while minimizing sequence gaps.

The isolated polynucleotide of the invention may be operably linked to an expression control sequence such as the pMT2 or pED expression vectors disclosed in Kaufman *et al.*, Nucleic Acids Res. 19, 4485-4490 (1991), in order to produce the protein recombinantly. Many suitable expression control sequences are known in the art. General methods of expressing recombinant proteins are also known and are exemplified in R. Kaufman, Methods in Enzymology 185, 537-566 (1990). As defined herein "operably linked" means that the isolated polynucleotide of the invention and an expression control sequence are situated within a vector or cell in such a way that the protein is expressed by a host cell which has been transformed (transfected) with the ligated polynucleotide/expression control sequence.

A number of types of cells may act as suitable host cells for expression of the protein. Mammalian host cells include, for example, monkey COS cells, Chinese Hamster Ovary (CHO) cells, human kidney 293 cells, human epidermal A431 cells,

human Colo205 cells, 3T3 cells, CV-1 cells, other transformed primate cell lines, normal diploid cells, cell strains derived from in vitro culture of primary tissue, primary explants, HeLa cells, mouse L cells, BHK, HL-60, U937, HaK or Jurkat cells.

Alternatively, it may be possible to produce the protein in lower eukaryotes such as yeast or in prokaryotes such as bacteria. Potentially suitable yeast strains include *Saccharomyces cerevisiae*, *Schizosaccharomyces pombe*, *Kluyveromyces* strains, *Candida*, or any yeast strain capable of expressing heterologous proteins. Potentially suitable bacterial strains include *Escherichia coli*, *Bacillus subtilis*, *Salmonella typhimurium*, or any bacterial strain capable of expressing heterologous proteins. If the protein is made in yeast or bacteria, it may be necessary to modify the protein produced therein, for example by phosphorylation or glycosylation of the appropriate sites, in order to obtain the functional protein. Such covalent attachments may be accomplished using known chemical or enzymatic methods.

The protein may also be produced by operably linking the isolated polynucleotide of the invention to suitable control sequences in one or more insect expression vectors, and employing an insect expression system. Materials and methods for baculovirus/insect cell expression systems are commercially available in kit form from, e.g., Invitrogen, San Diego, California, U.S.A. (the MaxBac® kit), and such methods are well known in the art, as described in Summers and Smith, Texas Agricultural Experiment Station Bulletin No. 1555 (1987), incorporated herein by reference. As used herein, an insect cell capable of expressing a polynucleotide of the present invention is "transformed."

The protein of the invention may be prepared by culturing transformed host cells under culture conditions suitable to express the recombinant protein. The resulting expressed protein may then be purified from such culture (i.e., from culture medium or cell extracts) using known purification processes, such as gel filtration and ion exchange chromatography. The purification of the protein may also include an affinity column containing agents which will bind to the protein; one or more column steps over such affinity resins as concanavalin A-agarose, heparin-toyopearl® or Cibacrom blue 3GA Sepharose®; one or more steps involving hydrophobic interaction chromatography using such resins as phenyl ether, butyl ether, or propyl ether; or immunoaffinity chromatography.

Alternatively, the protein of the invention may also be expressed in a form which will facilitate purification. For example, it may be expressed as a fusion protein, such as those of maltose binding protein (MBP), glutathione-S-transferase (GST) or thioredoxin (TRX). Kits for expression and purification of such fusion proteins are commercially available from New England BioLab (Beverly, MA), Pharmacia (Piscataway, NJ) and InVitrogen, respectively. The protein can also be tagged with an

epitope and subsequently purified by using a specific antibody directed to such epitope. One such epitope ("Flag") is commercially available from Kodak (New Haven, CT).

Finally, one or more reverse-phase high performance liquid chromatography (RP-HPLC) steps employing hydrophobic RP-HPLC media, e.g., silica gel having pendant methyl or other aliphatic groups, can be employed to further purify the protein. Some or all of the foregoing purification steps, in various combinations, can also be employed to provide a substantially homogeneous isolated recombinant protein. The protein thus purified is substantially free of other mammalian proteins and is defined in accordance with the present invention as an "isolated protein."

The protein of the invention may also be expressed as a product of transgenic animals, e.g., as a component of the milk of transgenic cows, goats, pigs, or sheep which are characterized by somatic or germ cells containing a nucleotide sequence encoding the protein.

The protein may also be produced by known conventional chemical synthesis. Methods for constructing the proteins of the present invention by synthetic means are known to those skilled in the art. The synthetically-constructed protein sequences, by virtue of sharing primary, secondary or tertiary structural and/or conformational characteristics with proteins may possess biological properties in common therewith, including protein activity. Thus, they may be employed as biologically active or immunological substitutes for natural, purified proteins in screening of therapeutic compounds and in immunological processes for the development of antibodies.

The proteins provided herein also include proteins characterized by amino acid sequences similar to those of purified proteins but into which modification are naturally provided or deliberately engineered. For example, modifications in the peptide or DNA sequences can be made by those skilled in the art using known techniques. Modifications of interest in the protein sequences may include the alteration, substitution, replacement, insertion or deletion of a selected amino acid residue in the coding sequence. For example, one or more of the cysteine residues may be deleted or replaced with another amino acid to alter the conformation of the molecule. Techniques for such alteration, substitution, replacement, insertion or deletion are well known to those skilled in the art (see, e.g., U.S. Patent No. 4,518,584). Preferably, such alteration, substitution, replacement, insertion or deletion retains the desired activity of the protein.

Other fragments and derivatives of the sequences of proteins which would be expected to retain protein activity in whole or in part and may thus be useful for screening or other immunological methodologies may also be easily made by those

skilled in the art given the disclosures herein. Such modifications are believed to be encompassed by the present invention.

### USES AND BIOLOGICAL ACTIVITY

5       The polynucleotides and proteins of the present invention are expected to exhibit one or more of the uses or biological activities (including those associated with assays cited herein) identified below. Uses or activities described for proteins of the present invention may be provided by administration or use of such proteins or by administration or use of polynucleotides encoding such proteins (such as, for example,  
10   in gene therapies or vectors suitable for introduction of DNA).

#### Research Uses and Utilities

15       The polynucleotides provided by the present invention can be used by the research community for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to  
20   compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; as a probe to "subtract-out" known sequences in the process of discovering other novel polynucleotides; for selecting and making oligomers for attachment to a "gene chip" or  
25   other support, including for examination of expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the polynucleotide can also be used in interaction trap  
30   assays (such as, for example, those described in Gyuris *et al.*, 1993, *Cell* 75: 791-803 and in Rossi *et al.*, 1997, *Proc. Natl. Acad. Sci. USA* 94: 8405-8410, all of which are incorporated by reference herein) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

35       The proteins provided by the present invention can similarly be used in assay to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which

the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation "Molecular Cloning: A Laboratory Manual", 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology: Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

#### Nutritional Uses

Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

#### Cytokine and Cell Proliferation/Differentiation Activity

A protein of the present invention may exhibit cytokine, cell proliferation (either inducing or inhibiting) or cell differentiation (either inducing or inhibiting) activity or may induce production of other cytokines in certain cell populations. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10,

B9, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7e and CMK.

The activity of a protein of the invention may, among other means, be measured by the following methods:

- 5        Assays for T-cell or thymocyte proliferation include without limitation those described in: *Current Protocols in Immunology*, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., *J. Immunol.* 137:3494-3500, 10    1986; Bertagnolli et al., *J. Immunol.* 145:1706-1712, 1990; Bertagnolli et al., *Cellular Immunology* 133:327-341, 1991; Bertagnolli, et al., *J. Immunol.* 149:3778-3783, 1992; Bowman et al., *J. Immunol.* 152: 1756-1761, 1994.

- Assays for cytokine production and/or proliferation of spleen cells, lymph node cells or thymocytes include, without limitation, those described in: Polyclonal T cell 15    stimulation, Kruisbeek, A.M. and Shevach, E.M. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 3.12.1-3.12.14, John Wiley and Sons, Toronto. 1994; and Measurement of mouse and human Interferon  $\gamma$ , Schreiber, R.D. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto. 1994.

- 20        Assays for proliferation and differentiation of hematopoietic and lymphopoietic cells include, without limitation, those described in: Measurement of Human and Murine Interleukin 2 and Interleukin 4, Bottomly, K., Davis, L.S. and Lipsky, P.E. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 6.3.1-6.3.12, John Wiley and Sons, Toronto. 1991; deVries et al., *J. Exp. Med.* 173:1205-1211, 1991; Moreau et al., 25    *Nature* 336:690-692, 1988; Greenberger et al., *Proc. Natl. Acad. Sci. U.S.A.* 80:2931-2938, 1983; Measurement of mouse and human interleukin 6 - Nordan, R. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 6.6.1-6.6.5, John Wiley and Sons, Toronto. 1991; Smith et al., *Proc. Natl. Acad. Sci. U.S.A.* 83:1857-1861, 1986; Measurement of human Interleukin 11 - Bennett, F., Giannotti, J., Clark, S.C. and Turner, K. J. In *Current* 30    *Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 6.15.1 John Wiley and Sons, Toronto. 1991; Measurement of mouse and human Interleukin 9 - Ciarletta, A., Giannotti, J., Clark, S.C. and Turner, K.J. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto. 1991.

- Assays for T-cell clone responses to antigens (which will identify, among others, 35    proteins that affect APC-T cell interactions as well as direct T-cell effects by measuring proliferation and cytokine production) include, without limitation, those described in: *Current Protocols in Immunology*, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience



(Chapter 3, In Vitro assays for Mouse Lymphocyte Function; Chapter 6, Cytokines and their cellular receptors; Chapter 7, Immunologic studies in Humans); Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

#### Immune Stimulating or Suppressing Activity

A protein of the present invention may also exhibit immune stimulating or immune suppressing activity, including without limitation the activities for which assays are described herein. A protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More specifically, infectious diseases caused by viral, bacterial, fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., malaria spp. and various fungal infections such as candidiasis. Of course, in this regard, a protein of the present invention may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein of the present invention include, for example, connective tissue disease, multiple sclerosis, systemic lupus erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitus, myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein of the present invention may also be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

Using the proteins of the invention it may also be possible to immune responses, in a number of ways. Down regulation may be in the form of inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of activated T cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, which involves

inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions (such as , for example, B7)), *e.g.*, preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (*e.g.*, B7- 1, B7-3) or blocking antibody), prior to transplantation can lead to the binding of the molecule to the natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant. Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins *in vivo* as described in Lenschow *et al.*, *Science* 257:789-792 (1992) and Turka *et al.*, *Proc. Natl. Acad. Sci USA*, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed., *Fundamental Immunology*, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function *in vivo* on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate

activation of T cells that are reactive against self tissue and which promote the production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by  
5 disrupting receptor:ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating  
10 autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/*lpr/lpr* mice or NZB hybrid mice, murine autoimmune collagen arthritis, diabetes mellitus in NOD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental  
15 Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example,  
20 enhancing an immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory forms of B lymphocyte antigens systemically.

Alternatively, anti-viral immune responses may be enhanced in an infected  
25 patient by removing T cells from the patient, costimulating the T cells *in vitro* with viral antigen- pulsed APCs either expressing a peptide of the present invention or together with a stimulatory form of a soluble peptide of the present invention and reintroducing the *in vitro* activated T cells into the patient. Another method of enhancing anti-viral immune responses would be to isolate infected cells from a patient, transfect them with  
30 a nucleic acid encoding a protein of the present invention as described herein such that the cells express all or a portion of the protein on their surface, and reintroduce the transfected cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to, and thereby activate, T cells *in vivo*.

In another application, up regulation or enhancement of antigen function  
35 (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (*e.g.*, sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide of the present invention can be administered to a subject to overcome tumor-

specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected *ex vivo* with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection *in vivo*.

The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules, can be transfected with nucleic acid encoding all or a portion of (*e.g.*, a cytoplasmic-domain truncated portion) of an MHC class I  $\alpha$  chain protein and  $\beta_2$  microglobulin protein or an MHC class II  $\alpha$  chain protein and an MHC class II  $\beta$  chain protein to thereby express MHC class I or MHC class II proteins on the cell surface. Expression of the appropriate class I or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (*e.g.*, B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for thymocyte or splenocyte cytotoxicity include, without limitation, those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500,

1986; Bowman et al., J. Virology 61:1992-1998; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

Assays for T-cell-dependent immunoglobulin responses and isotype switching  
 5 (which will identify, among others, proteins that modulate T-cell dependent antibody responses and that affect Th1/Th2 profiles) include, without limitation, those described in: Maliszewski, J. Immunol. 144:3028-3033, 1990; and Assays for B cell function: *In vitro* antibody production, Mond, J.J. and Brunswick, M. In *Current Protocols in Immunology*. J.E.e.a. Coligan eds. Vol 1 pp. 3.8.1-3.8.16, John Wiley and Sons, Toronto. 1994.

10 Mixed lymphocyte reaction (MLR) assays (which will identify, among others, proteins that generate predominantly Th1 and CTL responses) include, without limitation, those described in: *Current Protocols in Immunology*, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, *In Vitro* assays for Mouse Lymphocyte  
 15 Function 3.1-3.19; Chapter 7, *Immunologic studies in Humans*); Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

Dendritic cell-dependent assays (which will identify, among others, proteins expressed by dendritic cells that activate naive T-cells) include, without limitation,  
 20 those described in: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., *Journal of Experimental Medicine* 173:549-559, 1991; Macatonia et al., *Journal of Immunology* 154:5071-5079, 1995; Porgador et al., *Journal of Experimental Medicine* 182:255-260, 1995; Nair et al., *Journal of Virology* 67:4062-4069, 1993; Huang et al., *Science* 264:961-965, 1994; Macatonia et al., *Journal of Experimental Medicine* 169:1255-1264, 1989;  
 25 Bhardwaj et al., *Journal of Clinical Investigation* 94:797-807, 1994; and Inaba et al., *Journal of Experimental Medicine* 172:631-640, 1990.

Assays for lymphocyte survival/apoptosis (which will identify, among others, proteins that prevent apoptosis after superantigen induction and proteins that regulate lymphocyte homeostasis) include, without limitation, those described in:  
 30 Darzynkiewicz et al., *Cytometry* 13:795-808, 1992; Gorczyca et al., *Leukemia* 7:659-670, 1993; Gorczyca et al., *Cancer Research* 53:1945-1951, 1993; Itoh et al., *Cell* 66:233-243, 1991; Zacharchuk, *Journal of Immunology* 145:4037-4045, 1990; Zamai et al., *Cytometry* 14:891-897, 1993; Gorczyca et al., *International Journal of Oncology* 1:639-648, 1992.

Assays for proteins that influence early steps of T-cell commitment and  
 35 development include, without limitation, those described in: Antica et al., *Blood* 84:111-117, 1994; Fine et al., *Cellular Immunology* 155:111-122, 1994; Galy et al., *Blood* 85:2770-2778, 1995; Toki et al., *Proc. Nat. Acad Sci. USA* 88:7548-7551, 1991.

Hematopoiesis Regulating Activity

A protein of the present invention may be useful in regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells; in supporting the growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting the growth and proliferation of megakaryocytes and consequently of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation of hematopoietic stem cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantation, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either *in-vivo* or *ex-vivo* (i.e., in conjunction with bone marrow transplantation or with peripheral progenitor cell transplantation (homologous or heterologous)) as normal cells or genetically manipulated for gene therapy.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for proliferation and differentiation of various hematopoietic lines are cited above.

Assays for embryonic stem cell differentiation (which will identify, among others, proteins that influence embryonic differentiation hematopoiesis) include, without limitation, those described in: Johansson et al. Cellular Biology 15:141-151, 1995; Keller et al., Molecular and Cellular Biology 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

Assays for stem cell survival and differentiation (which will identify, among others, proteins that regulate lympho-hematopoiesis) include, without limitation, those described in: Methylcellulose colony forming assays, Freshney, M.G. In *Culture of Hematopoietic Cells*. R.I. Freshney, et al. eds. Vol pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; Primitive hematopoietic colony forming cells with high proliferative potential, McNiece, I.K. and

Briddell, R.A. In *Culture of Hematopoietic Cells*. R.I. Freshney, *et al.* eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., *Experimental Hematology* 22:353-359, 1994; Cobblestone area forming cell assay, Ploemacher, R.E. In *Culture of Hematopoietic Cells*. R.I. Freshney, *et al.* eds. Vol pp. 1-21, Wiley-Liss, Inc., New York, NY. 1994; Long term bone marrow cultures in the presence of stromal cells, Spooncer, E., Dexter, M. and Allen, T. In *Culture of Hematopoietic Cells*. R.I. Freshney, *et al.* eds. Vol pp. 163-179, Wiley-Liss, Inc., New York, NY. 1994; Long term culture initiating cell assay, Sutherland, H.J. In *Culture of Hematopoietic Cells*. R.I. Freshney, *et al.* eds. Vol pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

#### Tissue Growth Activity

A protein of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. *De novo* bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or

ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by a composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or  
 5 ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions of the present invention may provide an environment to attract tendon- or ligament-forming cells, stimulate growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of tendon/ligament  
 10 cells or progenitors *ex vivo* for return *in vivo* to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The protein of the present invention may also be useful for proliferation of  
 15 neural cells and for regeneration of nerve and brain tissue, *i.e.* for the treatment of central and peripheral nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral nerve injuries, peripheral  
 20 neuropathy and localized neuropathies, and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders, such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral  
 25 neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

30 It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition  
 35 or modulation of fibrotic scarring to allow normal tissue to regenerate. A protein of the invention may also exhibit angiogenic activity.



A protein of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein of the present invention may also be useful for promoting or  
 5 inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for tissue generation activity include, without limitation, those described  
 10 in: International Patent Publication No. WO95/16035 (bone, cartilage, tendon); International Patent Publication No. WO95/05846 (nerve, neuronal); International Patent Publication No. WO91/07491 (skin, endothelium ).

Assays for wound healing activity include, without limitation, those described  
 15 in: Winter, Epidermal Wound Healing, pps. 71-112 (Maibach, HI and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

#### Activin/Inhibin Activity

A protein of the present invention may also exhibit activin- or inhibin-related  
 20 activities. Inhibins are characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins and are characterized by their ability to stimulate the release of follicle stimulating hormone (FSH). Thus, a protein of the present invention, alone or in heterodimers with a member of the inhibin  $\alpha$  family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in  
 25 female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin-  $\beta$  group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from  
 30 cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein of the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

The activity of a protein of the invention may, among other means, be measured  
 35 by the following methods:

Assays for activin/inhibin activity include, without limitation, those described  
 in: Vale et al., Endocrinology 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986;

Vale et al., Nature 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986.

#### Chemotactic/Chemokinetic Activity

5 A protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or  
10 chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it  
15 can stimulate, directly or indirectly, the directed orientation or movement of such cell population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

20 The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis) consist of assays that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the  
25 adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W. Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12, Measurement of alpha and beta Chemokines 6.12.1-6.12.28; Taub et al. J.  
30 Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al. Eur. J. Immunol. 25: 1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol. 153: 1762-1768, 1994.

#### Hemostatic and Thrombolytic Activity

35 A protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulation disorders (including hereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma,

surgery or other causes. A protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system vessels (e.g., stroke).

5       The activity of a protein of the invention may, among other means, be measured by the following methods:

Assay for hemostatic and thrombolytic activity include, without limitation, those described in: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res. 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991); Schaub, Prostaglandins 35:467-474, 1988.

#### Receptor/Ligand Activity

A protein of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments of receptors and ligands) may themselves be useful as inhibitors of receptor/ligand interactions.

25       The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for receptor-ligand activity include without limitation those described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W. Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 7.28, Measurement of Cellular Adhesion under static conditions 7.28.1-7.28.22), Takai et al., Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al., J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp. Med. 169:149-160 1989; Stoltenborg et al., J. Immunol. Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995.

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#### Anti-Inflammatory Activity

Proteins of the present invention may also exhibit anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells

involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

#### 15      Cadherin/Tumor Invasion Suppressor Activity

Cadherins are calcium-dependent adhesion molecules that appear to play major roles during development, particularly in defining specific cell types. Loss or alteration of normal cadherin expression can lead to changes in cell adhesion properties linked to tumor growth and metastasis. Cadherin malfunction is also implicated in other human diseases, such as pemphigus vulgaris and pemphigus foliaceus (auto-immune blistering skin diseases), Crohn's disease, and some developmental abnormalities.

The cadherin superfamily includes well over forty members, each with a distinct pattern of expression. All members of the superfamily have in common conserved extracellular repeats (cadherin domains), but structural differences are found in other parts of the molecule. The cadherin domains bind calcium to form their tertiary structure and thus calcium is required to mediate their adhesion. Only a few amino acids in the first cadherin domain provide the basis for homophilic adhesion; modification of this recognition site can change the specificity of a cadherin so that instead of recognizing only itself, the mutant molecule can now also bind to a different cadherin. In addition, some cadherins engage in heterophilic adhesion with other cadherins.

E-cadherin, one member of the cadherin superfamily, is expressed in epithelial cell types. Pathologically, if E-cadherin expression is lost in a tumor, the malignant cells become invasive and the cancer metastasizes. Transfection of cancer cell lines with polynucleotides expressing E-cadherin has reversed cancer-associated changes by returning altered cell shapes to normal, restoring cells' adhesiveness to each other and to their substrate, decreasing the cell growth rate, and drastically reducing anchorage-independent cell growth. Thus, reintroducing E-cadherin expression reverts

carcinomas to a less advanced stage. It is likely that other cadherins have the same invasion suppressor role in carcinomas derived from other tissue types. Therefore, proteins of the present invention with cadherin activity, and polynucleotides of the present invention encoding such proteins, can be used to treat cancer. Introducing such proteins or polynucleotides into cancer cells can reduce or eliminate the cancerous changes observed in these cells by providing normal cadherin expression.

Cancer cells have also been shown to express cadherins of a different tissue type than their origin, thus allowing these cells to invade and metastasize in a different tissue in the body. Proteins of the present invention with cadherin activity, and polynucleotides of the present invention encoding such proteins, can be substituted in these cells for the inappropriately expressed cadherins, restoring normal cell adhesive properties and reducing or eliminating the tendency of the cells to metastasize.

Additionally, proteins of the present invention with cadherin activity, and polynucleotides of the present invention encoding such proteins, can be used to generate antibodies recognizing and binding to cadherins. Such antibodies can be used to block the adhesion of inappropriately expressed tumor-cell cadherins, preventing the cells from forming a tumor elsewhere. Such an anti-cadherin antibody can also be used as a marker for the grade, pathological type, and prognosis of a cancer, i.e. the more progressed the cancer, the less cadherin expression there will be, and this decrease in cadherin expression can be detected by the use of a cadherin-binding antibody.

Fragments of proteins of the present invention with cadherin activity, preferably a polypeptide comprising a decapeptide of the cadherin recognition site, and polynucleotides of the present invention encoding such protein fragments, can also be used to block cadherin function by binding to cadherins and preventing them from binding in ways that produce undesirable effects. Additionally, fragments of proteins of the present invention with cadherin activity, preferably truncated soluble cadherin fragments which have been found to be stable in the circulation of cancer patients, and polynucleotides encoding such protein fragments, can be used to disturb proper cell-cell adhesion.

Assays for cadherin adhesive and invasive suppressor activity include, without limitation, those described in: Hortsch et al. J Biol Chem 270 (32): 18809-18817, 1995; Miyaki et al. Oncogene 11: 2547-2552, 1995; Ozawa et al. Cell 63: 1033-1038, 1990.

#### Tumor Inhibition Activity

In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue

or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth.

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#### Other Activities

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in bone form or shape); effecting biorhythms or circadian cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein.

#### 30 ADMINISTRATION AND DOSING

A protein of the present invention (from whatever source derived, including without limitation from recombinant and non-recombinant sources) may be used in a pharmaceutical composition when combined with a pharmaceutically acceptable carrier. Such a composition may also contain (in addition to protein and a carrier) diluents, fillers, salts, buffers, stabilizers, solubilizers, and other materials well known in the art. The term "pharmaceutically acceptable" means a non-toxic material that does not interfere with the effectiveness of the biological activity of the active ingredient(s). The characteristics of the carrier will depend on the route of administration. The

pharmaceutical composition of the invention may also contain cytokines, lymphokines, or other hematopoietic factors such as M-CSF, GM-CSF, TNF, IL-1, IL-2, IL-3, IL-4, IL-5, IL-6, IL-7, IL-8, IL-9, IL-10, IL-11, IL-12, IL-13, IL-14, IL-15, IFN, TNF0, TNF1, TNF2, G-CSF, Meg-CSF, thrombopoietin, stem cell factor, and erythropoietin. The pharmaceutical composition may further contain other agents which either enhance the activity of the protein or complement its activity or use in treatment. Such additional factors and/or agents may be included in the pharmaceutical composition to produce a synergistic effect with protein of the invention, or to minimize side effects. Conversely, protein of the present invention may be included in formulations of the particular cytokine, lymphokine, other hematopoietic factor, thrombolytic or anti-thrombotic factor, or anti-inflammatory agent to minimize side effects of the cytokine, lymphokine, other hematopoietic factor, thrombolytic or anti-thrombotic factor, or anti-inflammatory agent.

A protein of the present invention may be active in multimers (e.g., heterodimers or homodimers) or complexes with itself or other proteins. As a result, pharmaceutical compositions of the invention may comprise a protein of the invention in such multimeric or complexed form.

The pharmaceutical composition of the invention may be in the form of a complex of the protein(s) of present invention along with protein or peptide antigens. The protein and/or peptide antigen will deliver a stimulatory signal to both B and T lymphocytes. B lymphocytes will respond to antigen through their surface immunoglobulin receptor. T lymphocytes will respond to antigen through the T cell receptor (TCR) following presentation of the antigen by MHC proteins. MHC and structurally related proteins including those encoded by class I and class II MHC genes on host cells will serve to present the peptide antigen(s) to T lymphocytes. The antigen components could also be supplied as purified MHC-peptide complexes alone or with co-stimulatory molecules that can directly signal T cells. Alternatively antibodies able to bind surface immunoglobulin and other molecules on B cells as well as antibodies able to bind the TCR and other molecules on T cells can be combined with the pharmaceutical composition of the invention.

The pharmaceutical composition of the invention may be in the form of a liposome in which protein of the present invention is combined, in addition to other pharmaceutically acceptable carriers, with amphipathic agents such as lipids which exist in aggregated form as micelles, insoluble monolayers, liquid crystals, or lamellar layers in aqueous solution. Suitable lipids for liposomal formulation include, without limitation, monoglycerides, diglycerides, sulfatides, lysolecithin, phospholipids, saponin, bile acids, and the like. Preparation of such liposomal formulations is within the level of skill in the art, as disclosed, for example, in U.S. Patent No. 4,235,871; U.S.

Patent No. 4,501,728; U.S. Patent No. 4,837,028; and U.S. Patent No. 4,737,323, all of which are incorporated herein by reference.

As used herein, the term "therapeutically effective amount" means the total amount of each active component of the pharmaceutical composition or method that is sufficient to show a meaningful patient benefit, i.e., treatment, healing, prevention or amelioration of the relevant medical condition, or an increase in rate of treatment, healing, prevention or amelioration of such conditions. When applied to an individual active ingredient, administered alone, the term refers to that ingredient alone. When applied to a combination, the term refers to combined amounts of the active ingredients that result in the therapeutic effect, whether administered in combination, serially or simultaneously.

In practicing the method of treatment or use of the present invention, a therapeutically effective amount of protein of the present invention is administered to a mammal having a condition to be treated. Protein of the present invention may be administered in accordance with the method of the invention either alone or in combination with other therapies such as treatments employing cytokines, lymphokines or other hematopoietic factors. When co-administered with one or more cytokines, lymphokines or other hematopoietic factors, protein of the present invention may be administered either simultaneously with the cytokine(s), lymphokine(s), other hematopoietic factor(s), thrombolytic or anti-thrombotic factors, or sequentially. If administered sequentially, the attending physician will decide on the appropriate sequence of administering protein of the present invention in combination with cytokine(s), lymphokine(s), other hematopoietic factor(s), thrombolytic or anti-thrombotic factors.

Administration of protein of the present invention used in the pharmaceutical composition or to practice the method of the present invention can be carried out in a variety of conventional ways, such as oral ingestion, inhalation, topical application or cutaneous, subcutaneous, intraperitoneal, parenteral or intravenous injection. Intravenous administration to the patient is preferred.

When a therapeutically effective amount of protein of the present invention is administered orally, protein of the present invention will be in the form of a tablet, capsule, powder, solution or elixir. When administered in tablet form, the pharmaceutical composition of the invention may additionally contain a solid carrier such as a gelatin or an adjuvant. The tablet, capsule, and powder contain from about 5 to 95% protein of the present invention, and preferably from about 25 to 90% protein of the present invention. When administered in liquid form, a liquid carrier such as water, petroleum, oils of animal or plant origin such as peanut oil, mineral oil, soybean oil, or sesame oil, or synthetic oils may be added. The liquid form of the



pharmaceutical composition may further contain physiological saline solution, dextrose or other saccharide solution, or glycols such as ethylene glycol, propylene glycol or polyethylene glycol. When administered in liquid form, the pharmaceutical composition contains from about 0.5 to 90% by weight of protein of the present invention, and preferably from about 1 to 50% protein of the present invention.

When a therapeutically effective amount of protein of the present invention is administered by intravenous, cutaneous or subcutaneous injection, protein of the present invention will be in the form of a pyrogen-free, parenterally acceptable aqueous solution. The preparation of such parenterally acceptable protein solutions, having due regard to pH, isotonicity, stability, and the like, is within the skill in the art. A preferred pharmaceutical composition for intravenous, cutaneous, or subcutaneous injection should contain, in addition to protein of the present invention, an isotonic vehicle such as Sodium Chloride Injection, Ringer's Injection, Dextrose Injection, Dextrose and Sodium Chloride Injection, Lactated Ringer's Injection, or other vehicle as known in the art. The pharmaceutical composition of the present invention may also contain stabilizers, preservatives, buffers, antioxidants, or other additives known to those of skill in the art.

The amount of protein of the present invention in the pharmaceutical composition of the present invention will depend upon the nature and severity of the condition being treated, and on the nature of prior treatments which the patient has undergone. Ultimately, the attending physician will decide the amount of protein of the present invention with which to treat each individual patient. Initially, the attending physician will administer low doses of protein of the present invention and observe the patient's response. Larger doses of protein of the present invention may be administered until the optimal therapeutic effect is obtained for the patient, and at that point the dosage is not increased further. It is contemplated that the various pharmaceutical compositions used to practice the method of the present invention should contain about 0.01  $\mu$ g to about 100 mg (preferably about 0.1mg to about 10 mg, more preferably about 0.1  $\mu$ g to about 1 mg) of protein of the present invention per kg body weight.

The duration of intravenous therapy using the pharmaceutical composition of the present invention will vary, depending on the severity of the disease being treated and the condition and potential idiosyncratic response of each individual patient. It is contemplated that the duration of each application of the protein of the present invention will be in the range of 12 to 24 hours of continuous intravenous administration. Ultimately the attending physician will decide on the appropriate duration of intravenous therapy using the pharmaceutical composition of the present invention.

Protein of the invention may also be used to immunize animals to obtain polyclonal and monoclonal antibodies which specifically react with the protein. Such antibodies may be obtained using either the entire protein or fragments thereof as an immunogen. The peptide immunogens additionally may contain a cysteine residue at the carboxyl terminus, and are conjugated to a hapten such as keyhole limpet hemocyanin (KLH). Methods for synthesizing such peptides are known in the art, for example, as in R.P. Merrifield, J. Amer.Chem.Soc. 85, 2149-2154 (1963); J.L. Krstenansky, *et al.*, FEBS Lett. 211, 10 (1987). Monoclonal antibodies binding to the protein of the invention may be useful diagnostic agents for the immunodetection of the protein. Neutralizing monoclonal antibodies binding to the protein may also be useful therapeutics for both conditions associated with the protein and also in the treatment of some forms of cancer where abnormal expression of the protein is involved. In the case of cancerous cells or leukemic cells, neutralizing monoclonal antibodies against the protein may be useful in detecting and preventing the metastatic spread of the cancerous cells, which may be mediated by the protein.

For compositions of the present invention which are useful for bone, cartilage, tendon or ligament regeneration, the therapeutic method includes administering the composition topically, systematically, or locally as an implant or device. When administered, the therapeutic composition for use in this invention is, of course, in a pyrogen-free, physiologically acceptable form. Further, the composition may desirably be encapsulated or injected in a viscous form for delivery to the site of bone, cartilage or tissue damage. Topical administration may be suitable for wound healing and tissue repair. Therapeutically useful agents other than a protein of the invention which may also optionally be included in the composition as described above, may alternatively or additionally, be administered simultaneously or sequentially with the composition in the methods of the invention. Preferably for bone and/or cartilage formation, the composition would include a matrix capable of delivering the protein-containing composition to the site of bone and/or cartilage damage, providing a structure for the developing bone and cartilage and optimally capable of being resorbed into the body. Such matrices may be formed of materials presently in use for other implanted medical applications.

The choice of matrix material is based on biocompatibility, biodegradability, mechanical properties, cosmetic appearance and interface properties. The particular application of the compositions will define the appropriate formulation. Potential matrices for the compositions may be biodegradable and chemically defined calcium sulfate, tricalciumphosphate, hydroxyapatite, polylactic acid, polyglycolic acid and polyanhydrides. Other potential materials are biodegradable and biologically well-defined, such as bone or dermal collagen. Further matrices are comprised of pure

proteins or extracellular matrix components. Other potential matrices are nonbiodegradable and chemically defined, such as sintered hydroxapatite, bioglass, aluminates, or other ceramics. Matrices may be comprised of combinations of any of the above mentioned types of material, such as polylactic acid and hydroxyapatite or collagen and tricalciumphosphate. The bioceramics may be altered in composition, such as in calcium- aluminate-phosphate and processing to alter pore size, particle size, particle shape, and biodegradability.

Presently preferred is a 50:50 (mole weight) copolymer of lactic acid and glycolic acid in the form of porous particles having diameters ranging from 150 to 800 microns. In some applications, it will be useful to utilize a sequestering agent, such as carboxymethyl cellulose or autologous blood clot, to prevent the protein compositions from disassociating from the matrix.

A preferred family of sequestering agents is cellulosic materials such as alkylcelluloses (including hydroxyalkylcelluloses), including methylcellulose, ethylcellulose, hydroxyethylcellulose, hydroxypropylcellulose, hydroxypropylmethylcellulose, and carboxymethylcellulose, the most preferred being cationic salts of carboxymethylcellulose (CMC). Other preferred sequestering agents include hyaluronic acid, sodium alginate, poly(ethylene glycol), polyoxyethylene oxide, carboxyvinyl polymer and poly(vinyl alcohol). The amount of sequestering agent useful herein is 0.5-20 wt%, preferably 1-10 wt% based on total formulation weight, which represents the amount necessary to prevent desorption of the protein from the polymer matrix and to provide appropriate handling of the composition, yet not so much that the progenitor cells are prevented from infiltrating the matrix, thereby providing the protein the opportunity to assist the osteogenic activity of the progenitor cells.

In further compositions, proteins of the invention may be combined with other agents beneficial to the treatment of the bone and/or cartilage defect, wound, or tissue in question. These agents include various growth factors such as epidermal growth factor (EGF), platelet derived growth factor (PDGF), transforming growth factors (TGF- $\alpha$  and TGF- $\beta$ ), and insulin-like growth factor (IGF).

The therapeutic compositions are also presently valuable for veterinary applications. Particularly domestic animals and thoroughbred horses, in addition to humans, are desired patients for such treatment with proteins of the present invention.

The dosage regimen of a protein-containing pharmaceutical composition to be used in tissue regeneration will be determined by the attending physician considering various factors which modify the action of the proteins, e.g., amount of tissue weight desired to be formed, the site of damage, the condition of the damaged tissue, the size of a wound, type of damaged tissue (e.g., bone), the patient's age, sex, and diet, the

severity of any infection, time of administration and other clinical factors. The dosage may vary with the type of matrix used in the reconstitution and with inclusion of other proteins in the pharmaceutical composition. For example, the addition of other known growth factors, such as IGF I (insulin like growth factor I), to the final composition, may  
5 also effect the dosage. Progress can be monitored by periodic assessment of tissue/bone growth and/or repair, for example, X-rays, histomorphometric determinations and tetracycline labeling.

Polynucleotides of the present invention can also be used for gene therapy. Such polynucleotides can be introduced either *in vivo* or *ex vivo* into cells for expression  
10 in a mammalian subject. Polynucleotides of the invention may also be administered by other known methods for introduction of nucleic acid into a cell or organism (including, without limitation, in the form of viral vectors or naked DNA).

Cells may also be cultured *ex vivo* in the presence of proteins of the present invention in order to proliferate or to produce a desired effect on or activity in such  
15 cells. Treated cells can then be introduced *in vivo* for therapeutic purposes.

Patent and literature references cited herein are incorporated by reference as if fully set forth.

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